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EXPRESS MAIL LABEL NO. EL828141124US DATE OF DEPOSIT: May 1, 2001

FORM PTO-	1390		U.S. DEPARTMENT OF C	COMMERCE PATENT AND TRADEMARK OFFICE	ATTORNEY'S DOCKET NUMBER									
	TR	ΔΝ	SMITTAL LETTER	178-59010										
			SIGNATED/ELECTE											
			ICERNING A FILING	U.S. APPLICATION NO. (If known, see 37 C.F.R. § 1.5)										
			·	09/831000										
PCT/US			APPLICATION NO.	PRIORITY DATE CLAIMED										
				November 5, 1999	November 6, 1998									
TITLE OF INVENTION CLONING OF RHESUS MACAQUE RHADINOVIRUS GENOME AND METHODS FOR ITS USE														
[APPLICA]	APPLICANT(S) FOR DO/EO/US													
Scott W. Wong, Michael K. Axthelm, and Robert P. Searles Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:														
1. This is a FIRST submission of items concerning a filing under 35 U.S.C. § 371.														
	2.													
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ر. د	3. This is an express request to begin national examination procedures (35 U.S.C. § 371(f) at any time rather than delay examination until the expiration of the applicable time limit set in 35 U.S.C. § 371(b) and PCT Articles 22 and 39(1).													
4. A proper Demand for International Preliminary Examination was made by the 19 th month from the earliest clapriority date.														
	5.	\boxtimes	A copy of the International A	pplication as filed (35 U.S.C. § 371(c)(2))										
		a. is transmitted herewith (required only if not transmitted by the International Bureau).												
			b. has been transmitted by											
		c. is not required, as the application was filed in the United States Receiving Office (RO/US).												
	6.													
	7.	\boxtimes												
		a. are transmitted herewith (required only if not transmitted by the International Bureau).												
		b. have been transmitted by the International Bureau.												
		c. have not been made; however, the time limit for making such amendments has NOT expired.												
	d. \(\text{\tin\text{\texi}\text{\text{\text{\text{\text{\text{\text{\text{\text{\texi}\text{\text{\text{\text{\t													
	8. A translation of the amendments to the claims under PCT Article 19 (35 U.S.C. § 371(c)(3)).													
	9. An oath or declaration of the inventor(s) (35 U.S.C. § 371(c)(4)).													
	10. A translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. § 371(c)(5)).													
	Items 11. to 16. below concern document(s) or information included: 11. An Information Disclosure Statement under 37 C.F.R. §§ 1.97 and 1.98.													
	 12. An assignment document for recording. A separate cover sheet in compliance with 37 C.F.R. §§ 3.28 and 3.31 and the Recordal fee of \$40.00 is included. 													
	13. A FIRST preliminary amendment.													
	☐ A SECOND or SUBSEQUENT preliminary amendment.													
	14. ⊠ Sequence Listing: Paper copy, <u>293</u> pages; computer readable copy on diskette.													
	15. Statement in Compliance with 37 C.F.R. § 1.821(f) verifying identity of above copies.													
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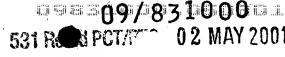
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EXPRESS MAIL LABEL NO. EL828141124US DATE OF DEPOSIT: May 1, 2001

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d. 🛛	Please return the enclosed postcard to confirm that the items listed above have been received.										
NOTE:	Where an appropriate time limit under 37 C.F.R. § 1.494 or § 1.495 has not been met, a petition to revive (37 C.F.R. § 1.137(a or (b)) must be filed and granted to restore the application to pending status.										
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DATE OF DEPOSIT: May 2, 2001
Attorney Reference Number 178-59010
Application Number

#4A

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: Wong et al.

Application No.: To be assigned

Filed: Herewith

For: CLONING OF RHESUS MACAQUE

RHADINOVIRUS GENOME AND METHODS

FOR ITS USE

Examiner: To be assigned

Date: May 2, 2001

Art Unit: To be assigned

CERTIFICATE OF EXPRESS MAILING

I hereby certify that this paper and the documents referred to as being attached or enclosed herewith are being deposited with the United States Postal Service on May 2, 2001 as Express Mail Label No. EL828141124US in an envelope addressed to: BOX PCT, COMMISSIONER FOR PATENTS, WASHINGTON, D.C. 20231.

William D. Noonan, M.D. Attorney for Applicant

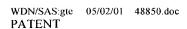
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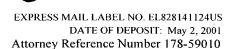
PRELIMINARY AMENDMENT

Prior to examination of the above-identified application, please amend the claims as follows:

Please cancel claims 1-34, and insert the following new claims:

- 35. (New) An isolated virus (RRV) as deposited with ATCC as deposit accession number VR-2601.
 - 36. (New) A purified virus, having a nucleic acid sequence
 - (a) shown in SEQ ID NO:1 or
 - (b) a conservative variant thereof.
- 37. (New) The purified virus of claim 36, wherein the nucleic acid sequence has at least 95% sequence identity to the nucleic acid sequence shown in SEQ ID NO:1.



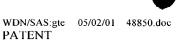


Application Number

38. (New) A purified protein encoded by an open reading frame of the virus of claim 36.

cont

- (New) A purified protein of claim 38, wherein the protein comprises an amino acid sequence selected from the group consisting of:
 - (a) an amino acid sequence shown in odd numbered sequences of SEQ ID NOS:3-165; and
 - (b) amino acid sequences that differ from those specified in (a) by one or more conservative amino acid substitutions wherein the function of the protein is preserved.
- 40. (New) A purified protein with an amino acid sequence that is at least 95% sequence identity to the sequences specified in claim 39.
- 41. (New) The purified protein of claim 39, wherein the amino acid sequence is selected from odd numbered sequences within the group consisting of SEQ ID NOS:3-19 and 23-165.
- 42. (New) An isolated nucleic acid molecule encoding a protein according to claim 39.
- 43. (New) An isolated nucleic acid molecule according to claim 42, wherein the molecule comprises a sequence selected from the group consisting of even numbered sequences of SEQ ID NOS:2-164.



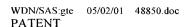


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DATE OF DEPOSIT: May 2, 2001
Attorney Reference Number 178-59010
Application Number

44. (New) The isolated nucleic acid molecule according to claim 43, wherein the molecule comprises a sequence selected from the group consisting of even numbered sequences of SEQ ID NOS:2-18 and 22-164.



- 45. (New) A recombinant nucleic acid molecule comprising a promoter sequence operably linked to a nucleic acid molecule according to claim 42.
- 46. (New) A cell transformed with a recombinant nucleic acid molecule according to claim 42.
 - 47. (New) A non-human mammal purposefully infected with the virus of claim 36.
 - 48. (New) The mammal of claim 47, wherein the mammal is a primate.
- 49. (New) An oligonucleotide comprising a sequence selected from the group consisting of:
 - (a) at least 20 contiguous nucleotides of the nucleic acid sequence of the virus of claim 36;
- (b) at least 30 contiguous nucleotides of the nucleic acid sequence of the virus of claim 36; and
- (c) at least 50 contiguous nucleotides of the nucleic acid sequence of the virus of claim 36.
 - 50. (New) An isolated nucleic acid molecule that encodes the protein of claim 40.
 - 51. (New) An isolated nucleic acid molecule encoding a protein of claim 40.



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52. (New) An isolated nucleic acid molecule encoding all proteins encoded by the virus of claim 36, and having a biological activity of an RRV virus.

- 53. (New) A method for testing the efficacy of a drug in the treatment of a condition associated with the virus of claim 36, the method comprising:
 - (a) administering the drug to a non-human primate infected with the virus of claim 36; and
 - (b) observing the primate to determine if the drug prevents or reduces the presentation of one or more symptoms associated with viral infection.
 - 54. (New) The method of claim 53, wherein the primate is immunocompromised.
 - 55. (New) The method of claim 54, wherein the drug is for the treatment of Kaposi's sarcoma and lymphoproliferative disorders.
 - 56. (New) The method of claim 54, wherein the primate is immuno-compromised as a result of infection by Simian Immunodeficiency Virus (SIV).
 - 57. (New) The method of claim 53, wherein the condition associated with infection with the virus is one or more of B-cell hyperplasia, lymphadenopathy, splenomegaly, hypergammaglobinulinemia or autoimmune hemolytic anemia.
 - 58. (New) The method of claim 53, wherein the non-human primate is a Rhesus macaque monkey.





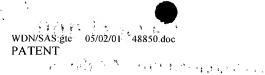
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- 59. (New) A method for producing a non-human primate model for testing potential treatments for a condition associated an infection with the virus of claim 36, comprising
- (a) administering a treatment to the primate to render the primate immunocompromised; and
 - (b) infecting the primate with the virus of claim 36.
- 60. (New) The method of claim 59, wherein the condition is Kaposi's sarcoma and lymphoproliferative disorders.
- 61. (New) The method of claim 59, wherein the treatment used to render the primate immuno-compromised is infection with SIV.
- 62. (New) The method of claim 59, wherein the non-human primate is a Rhesus macaque monkey.
- 63. (New) A method for testing the efficacy of a candidate vaccine against the virus of claim 36, or conditions associated infection with the virus of claim 36, the method comprising:
- (a) administering the vaccine to a subject capable of infection with the virus of claim 36;
 - (b) inoculating the subject with the virus; and
- (c) observing the subject to determine if the vaccine prevents or reduces an incidence of viral infection or presentation of one or more conditions associated with the viral infection.
 - 64. (New) The method of claim 63, wherein the subject is a primate.
 - 65. (New) The method of claim 64, wherein the primate is a non-human primate.





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Application Number

cont

- 66. (New) The method of claim 63, wherein the primate is immunocompromised.
- 67. (New) The method of claim 63, wherein the conditions associated with infection include B-cell hyperplasia, lymphadenopathy, splenomegaly, hypergammaglobinulinemia or autoimmune hemolytic anemia.
- 68. (New) The method of claim 65, wherein the non-human primate is a Rhesus macaque monkey.

CONCLUSION

No new matter is added. Entry of this amendment is respectfully requested prior to examination. If any minor matters remain to be addressed prior to examination, the Examiner is invited to call the undersigned at the telephone number listed below.

Respectfully submitted,

KLARQUIST SPARKMAN CAMPBELL LEIGH & WHINSTON, LLP

 $\mathbf{B}\mathbf{y}$

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DATE OF DEPOSIT: May 2, 2001 Attorney Reference Number 178-59010 Application Number

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: Wong et al.

WDN/SAS:gte 05/02/01 49081

PATENT

Application No.: To be assigned

Filed: Herewith

For: CLONING OF RHESUS MACAQUE

RHADINOVIRUS GENOME AND METHODS

FOR ITS USE

Examiner: To be assigned

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WASHINGTON, D.C. 20231.

William D. Noonan, M.D.
Attorney for Applicant

STATEMENT IN COMPLIANCE WITH 37 C.F.R. § 1.821(f)

BOX PCT COMMISSIONER FOR PATENTS Washington, DC 20231

Sir:

In compliance with 37 C.F.R. § 1.821(f), the undersigned declares that the nucleotide and/or amino acid sequences presented in the paper copy of the "Sequence Listing" submitted herewith are the same as the sequences contained in the computer-readable form of said "Sequence Listing." No new matter has been added.

Respectfully submitted,

KLARQUIST SPARKMAN CAMPBELL LEIGH & WHINSTON, LLP

y William D. Naan

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CLONING OF RHADINOVIRUS GENOME AND METHODS FOR ITS USE

FIELD OF THE INVENTION

The invention relates to the genome of a rhesus macaque rhadinovirus and provides compositions and methods for the production of animal models useful in assessing the efficacy of drugs and vaccines in the treatment and prevention of conditions associated with infection by the virus, such as Kaposi's sarcoma and lymphoproliferative disorders.

BACKGROUND

Converging lines of evidence indicate that Kaposi's sarcoma-associated herpesvirus (KSHV) is the etiological agent responsible for Kaposi's sarcoma (KS) in individuals with and without HIV infection (Chang et al., 1994, Science 266:1865-9; Schalling et al., 1995, Nature Med. 7:707-8; Moore & Chang, 1995, N. Engl. J. Med. 332:1181-5; Whitby et al., 1995, Lancet 346:799-802; Ambroziak et al., 1995, Science 268:582-3.; Dupin et al., 1995, Lancet 345:761-2.; Chuck et al., 1996, J. Infect. Dis. 173:248-51; O'Neill et al., 1996, J. Clin. Pathol. 49:306-8; Gao et al., 1996, Nature Med. 2:925-8; Kedes et al., 1996, Nature Med. 2:918-24; Gao et al., 1996, N. Engl. J. Med. 335:233-41). In addition to KS, KSHV is also responsible for other acquired immunodeficiency syndrome (AIDS)-related and non-AIDS-related malignancies, such as primary effusion lymphomas (Cesarman et al., 1995, N. Engl. J. Med. 332:1186-91; Nador et al., 1996, Blood 88:645-56; Otsuki et al, 1996, Leukemia 10:1358-62), and multicentric Castleman's disease (MCD), a B cell proliferation disorder associated with overexpression of IL-6 activity (Soulier et al., 1995, Blood 86:1276-80; Yoshizaki et al., 1989, Blood 74:1360-7).

More recently, KSHV has been proposed to be involved in multiple myeloma, a B cell abnormality of monoclonal origin (Rettig et al., 1997, *Science* 276:1851-4; Said et al., 1997, *Blood* 90:4278-82; Parravicini et al., 1997, *Science* 278:1969-70; Masood et al., 1997, *Science* 278:1970-1; Whitby et al., 1997, *Science* 278:1971-2; Cottoni et al., 1997, *Science* 278:1972; Brousset et al., 1997, *Science* 278:290-4). Understanding how KSHV is involved in these malignancies is important for the generation of therapies against the spectrum of KSHV-associated diseases.

Testing the efficacy of therapeutics and vaccines against any disease, such as KHSV, is greatly facilitated by the availability of an animal model, such as a non-human primate model, because non-human primates are physiologically very similar to humans. Although such models have been developed for the study of HIV infection (for example, U.S. Patent Nos. 5,212,084 and 5,543,131) none has yet been developed for KSHV infection.

Infection of animals with some herpesviruses, namely *Herpesvirus saimiri* and murine herpesvirus type 68, can cause a lymphoproliferative disorder (LPD). However, these animals are not adequate models of KSHV pathogenesis because they lack certain KSHV genes that may

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contribute to viral pathogenesis or influence HIV infection, such as Interleukin 6 (IL-6) and macrophage inflammatory protein 1 (MIP-1) (Albrecht et al., 1992, J. Virol. 66:5047-58; Virgin et al., 199% J. Virol. 71:5894-904). Thus, so far the establishment of a non-human primate model for KSHV infection has remained elusive.

The present invention addresses this problem, and others, in the development of animal models for a variety of pathological conditions and diseases.

SUMMARY OF THE DISCLOSURE

Rhesus macaques naturally harbor a virus related to KSHV, referred to as RRV, for rhesus rhadinovirus. Genetic analysis of RRV reveals the presence of an IL-6-like gene in a position analogous to that of the KSHV IL-6. The present disclosure also includes information about pathological conditions associated with RRV infection.

The present invention provides the genomic sequence (nucleotide and amino acid) for the RRV genome and its use for developing a non-human primate model for KSHV infection. The invention includes the genome of the newly isolated Rhesus macaque rhadinovirus, RRV isolate 17577 (referred to herein as RRV), but the invention includes variant RRV viruses and related viruses that infect other species. RRV shows some similarity to human Kaposi's sarcoma-associated herpes virus (KSHV, also called HHV8) and possesses genes for both IL-6 and MIP.

The invention encompasses the isolated polynucleotide genome of RRV as shown in SEQ ID NO 1, and the identified ORFs (open reading frames) of this genome (even-numbered SEQ ID NOS 2-164). Also included within the invention are oligonucleotides comprising at least 15, 20, 30, 40, 50, 70, 100 and 150 consecutive nucleotides of the genome sequence as shown in SEQ ID NO 1. Additionally, the invention encompasses various segments of the RRV genome as shown in SEQ ID NO 1, for instance, segments consisting of 999 nucleotides, for example, from nucleotide 1-999, 1000-1999, 2000-2999, 3000-3999, 4000-4999 and so on until the end of the nucleotide sequence. Proteins and parts of proteins encoded within such segments are also covered by the invention.

The invention also includes purified proteins encoded by the RRV genome, the amino acid sequences of which are shown in odd-numbered SEQ ID NOS 3-165. Proteins that have defined degrees of sequence identity with the proteins of SEQ ID NO 1 are also within the scope of the invention. Such proteins may display, for example, at least 50%, 55%, 60%, 70%, 80%, 90%, 95% or even 98% or greater amino acid sequence identity with the native proteins.

The invention further includes nucleic acids encoding the RRV proteins as well as recombinant nucleic acids that include a promoter operably linked to a nucleic acid that encodes an RRV protein.

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Additionally included are isolated nucleic acid molecules of various defined lengths that show at least 50%, 60%, 70%, 80%, 90%, 95%, 98% or 100% sequence identity with an RRV ORF sequence, such as the sequence shown in SEQ ID NO 1, or in one of the other sequence listings. The invention also includes isolated nucleic acid molecules of various defined lengths that hybridize with an ORF as shown in SEQ ID NO 1 under wash hybridization conditions of about 70°C and 0.2 x SSC for 1 hour, or alternatively under less stringent conditions of 65°C, 60°C, or 55°C with about 0.2 to 2 x SSC (with, for instance, about 0.1% SDS) for 1 hour.

Also within the invention are cells and virions that contain the nucleic acid molecules as described above.

Additionally the scope of the invention includes the nucleic acid sequences defined by nucleotides 1 to 11031 of SEQ ID NO 1 and nucleotides 21625 to 133719 of SEQ ID NO 1, and ORFs selected from these nucleic acid sequences. The invention also includes isolated nucleic acid molecules of various defined lengths that show at least 50%, 60%, 70%, 80%, 90%, 95% or 98% sequence identity with, an ORF contained within nucleotides 1-11031 or 21625-133719 of the nucleotide sequence as shown in SEQ ID NO 1. Alternatively, the invention includes at least 15, 20, 30, 40, 50, 70, 100 or 150 consecutive nucleotides within nucleotides 1 to 11031 of SEQ ID NO 1 and nucleotides 21625 to 133719 of SEQ ID NO 1, or within ORFs selected from these nucleic acid sequences.

Also included are isolated nucleic acid molecules of various lengths that hybridize under wash conditions of 70°C and about 0.2 x SSC for 1 hour, or alternatively under less stringent conditions of 65°C, 60°C, or 55°C with from about 0.2 to 2 x SSC (with, for instance, about 0.1% SDS) for 1 hour, with an ORF of nucleotides 1-11031 or 21625-133719 of the nucleotide sequence as shown in SEQ ID NO 1.

Recombinant molecules are also encompassed within the bounds of the invention, and include, for instance, a nucleic acid molecule encoding an RRV protein (or fragments or variants thereof) linked to a non-native nucleic acid sequence such as a promoter. The nucleic acid molecule linked to the promoter may be all or part of an ORF encoding an RRV protein, such as any ORF of SEQ ID NO 1, may be one or more fragments of a DNA sequence selected from the DNA sequence defined by nucleotides 1 to 11031 and nucleotides 21625 to 133719 as shown in SEQ ID NO 1, or DNA sequences encoding variants or fragments of proteins encoded by those sequences.

The present invention also relates to the isolation of a virus (RRV) from a rhesus macaque monkey which, when experimentally introduced into immuno-compromised macaques, produces pathological conditions, such as disease signs and symptoms, that parallel those seen in human subjects infected with KSHV, including lymphoproliferative disorders (LPD), lymphadenopathy, splenomegaly, B cell hyperplasia, autoimmune hemolytic anemia, retroperitoneal fibromatosis (a Kaposi's sarcoma-like mesenchymal proliferative disease of body cavities), and

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hypergammaglobulinemia, wherein the virus encodes homologues of IL-6 and MIP-1 which are similar to KSHV.

One aspect of the present invention is the isolated virus, RRV, and related species and other variants thereof. In another aspect of the invention, the virus is used to produce a non-human primate model for KSHV infection, or diseases associated with RRV infection; such a model may be produced, for example, by infecting a non-human primate (such as an immunocompromised non-human primate) with RRV. This model may thus be used to evaluate the efficacy of candidate therapeutics and vaccines for KSHV infection treatment and prophylaxis, or other pathological conditions associated with RRV infection. Although it is not required that the primate be first immuno-compromised and then infected with RRV, particular embodiments of the animal model include both infecting the primate with the virus and rendering it immuno-compromised (or equivalently obtaining an already immunocompromised primate).

In another embodiment, the invention provides a method for testing the efficacy of a drug in the treatment of Kaposi's sarcoma and lymphoproliferative disorders or other pathological conditions associated with RRV infection, by administering the drug to an immuno-compromised non-human primate infected with RRV, and then observing the primate to determine if the drug prevents or reduces the presentation of one or more signs, symptoms, laboratory abnormalities, or other pathological conditions associated with infection with the virus. Such conditions include B-cell hyperplasia, lymphadenopathy, splenomegaly, hypergammaglobinulinemia, retroperitoneal fibromatosis (a Kaposi's sarcoma-like mesenchymal proliferative disease of body cavities), and autoimmune hemolytic anemia. The efficacy of a vaccine to prevent KSHV infection, or pathological conditions associated with RRV infection, may similarly be assessed by administering the candidate vaccine to the animal and then attempting to infect the animal with RRV. In particular embodiments, the animal to which the candidate vaccine is administered may be an immunocompromised animal. Failure to infect the animal, when control animals not given the candidate vaccine do become infected, indicates that the vaccine conferred protection.

The foregoing and other objects, features, and advantages of the invention will become more apparent from the following detailed description of several examples which proceeds with reference to the accompanying figures.

BRIEF DESCRIPTION OF THE FIGURES

FIG. 1 shows a phylogenetic comparison of the gammaherpesviruses Epstein-Barr virus (EBV), Alcelaphine herpesvirus (AHV), Murine herpesvirus (MHV), Herpesvirus saimiri (HVS), Kaposi's sarcoma-associated herpesvirus (KSHV), and Rheusus rhadinovirus 17577 (RRV). It shows that among the known sequenced viruses, RRV is the closest relative to KSHV, using an accepted maximum parsimony method of evaluating evolutionary relationships.

FIG. 2 is a table showing the BamHI, EcoRI and HindIII restriction fragments of the RRV

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FIG. 3 is a schematic map of the ORFs of RRV. Arrow direction represents direction of transcription.

FIG. 4 is a table showing the size, location and description (similarity to other proteins) of the proteins encoded by the ORFs of RRV.

FIG. 5 is a table showing a comparison of corresponding repeats in RRV and KSHV.

FIG. 6 is a table showing the comparison of interferon regulatory elements encoded by RRV and KSHV.

FIG. 7 is a table comparing the ORFs of RRV, KSHV and HVS. The table shows the start and stop points, the strand (+ or -) from which the ORF is transcribed, the size of the ORFs and the percentage similarity of KSHV and HVS when compared with RRV.

FIGS. 8A-8D are graphs showing CD20+ lymphocytes, antibody response and RhKSHV isolation/detection in macaques infected with SIVmac239 and RRV (A)18483 and (B) 18570 and macaques infected with SIVmac239 only (C) 18503 and (D) 18540. A "+" indicates positive for virus culture or viral DNA, as defined by PCR and Southern blot analysis; "-", negative for virus culture or viral DNA.

FIG. 9 shows the result of the PCR analysis of PBLs and LNMCs from each of the macaques (18483, 18503, 18540 and 18570) for RRV DNA and β -globin in (A) graphical form and (B) digital form.

FIG. 10 shows the DNA sequence of the RRV ORF that encodes the IL-6 protein. The corresponding translated polypeptide sequence is shown in standard three letter code below the DNA sequence.

FIG. 11 shows the DNA sequence of the RRV ORF that encodes the MIP protein. The corresponding translated polypeptide sequence is shown in standard three letter code below the DNA sequence.

SEQUENCE LISTING

The nucleic and amino acid sequences listed in the accompanying sequence listing are shown using standard letter abbreviations for nucleotide bases, and the code for amino acids. Only one strand of each nucleic acid sequence is shown, but the complementary strand is understood as included by any reference to the displayed strand.

SEQ ID NO 1 shows the nucleotide sequence of the RRV genome.

SEQ ID NO 2 shows the cDNA nucleotide sequence of RRV R1, corresponding to nucleotides 1353-2624 of SEQ ID NO 1.

SEQ ID NO 3 shows the amino acid sequence of the RRV R1 protein.

SEQ ID NO 4 shows the cDNA nucleotide sequence of RRV ORF 2, corresponding to the complement of nucleotides 2692-3258 of SEQ ID NO 1, which encodes dihydrofolate reductase,

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and which has some similarity to Kaposi's sarcoma-associated herpesvirus (KSHV) ORF 2.

SEQ ID NO 5 shows the amino acid sequence of the ORF2 protein, dihydrofolate reductase protein, which has some similarity to KSHV ORF 2 protein.

SEQ ID NO 6 shows the cDNA nucleotide sequence of RRV ORF 4, corresponding to nucleotides 3676-5613 of SEQ ID NO 1, which encodes complement binding protein, and which has some similarity to KSHV ORF 4.

SEQ ID NO 7 shows the amino acid sequence of the RRV ORF 4 protein, complement binding protein, corresponding to nucleotides 6045-9443 of SEQ ID NO 1, and which has some similarity to KSHV ORF 4 protein.

SEQ ID NO 8 shows the cDNA nucleotide sequence of RRV ORF 6, corresponding to nucleotides 6045-9443 of SEQ ID NO 1, which encodes ssDNA binding protein, and which has some similarity to KSHV ORF 6.

SEQ ID NO 9 shows the amino acid sequence of the RRV ORF 6 protein, ssDNA binding protein, which has some similarity to KSHV ORF 6 protein.

SEQ ID NO 10 shows the cDNA nucleotide sequence of RRV ORF 7, corresponding to nucleotides 9468-11528 of SEQ ID NO 1, which encodes a transport protein, and which has some similarity to KSHV ORF 7.

SEQ ID NO 11 shows the amino acid sequence of the RRV ORF 7 protein, transport protein, which has some similarity to KSHV ORF 7 protein.

SEQ ID NO 12 shows the cDNA nucleotide sequence of RRV ORF 8, corresponding to nucleotides 11515-14004 of SEQ ID NO 1, which encodes glycoprotein B, and which has some similarity to KSHV ORF 8.

SEQ ID NO 13 shows the amino acid sequence of the RRV ORF 8 protein, glycoprotein B protein, which has some similarity to KSHV ORF 8 protein.

SEQ ID NO 14 shows the cDNA nucleotide sequence of RRV ORF 9, DNA polymerase protein, corresponding to nucleotides 14122-17166 of SEQ ID NO 1, which has some similarity to KSHV ORF 9.

SEQ ID NO 15 shows the amino acid sequence of the RRV ORF 9 protein, DNA polymerase protein, which has some similarity to KSHV ORF 9.

SEQ ID NO 16 shows the cDNA nucleotide sequence of RRV ORF 10, corresponding to nucleotides 17261-18511 of SEQ ID NO 1, which has some similarity to KSHV ORF 10.

SEQ ID NO 17 shows the amino acid sequence of the RRV ORF 10 protein, which has some similarity to KSHV ORF 10.

SEQ ID NO 18 shows the cDNA nucleotide sequence of RRV ORF 11, corresponding to nucleotides 18520-19749 of SEQ ID NO 1, which has some similarity to KSHV ORF 11.

SEQ ID NO 19 shows the amino acid sequence of the RRV ORF 11 protein, which has some similarity to KSHV ORF 11.

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SEQ ID NO 20 shows the cDNA nucleotide sequence of RRV R2, corresponding to the complement of nucleotides 19921-20544 of SEQ ID NO 1, which has some similarity to the Kaposi's sarcoma-associated IL-6 gene.

SEQ ID NO 21 shows the amino acid sequence of the RRV R2 protein which has some similarity to IL-6.

SEQ ID NO 22 shows the cDNA nucleotide sequence of RRV ORF 70, thymidylate synthase, corresponding to the complement of nucleotides 20777-21778 of SEQ ID NO 1, and which has some similarity to KSHV ORF 70.

SEQ ID NO 23 shows the amino acid sequence of the RRV ORF 70 protein, thymidylate synthase, which has some similarity to KSHV ORF 70 protein.

SEQ ID NO 24 shows the cDNA nucleotide sequence of RRV R3, corresponding to the complement of nucleotides 22245-22592 of SEQ ID NO 1, which has some similarity to the KSHV K4 viral MIP gene.

SEQ ID NO 25 shows the amino acid sequence of the RRV R3 protein, which has some similarity to KSHV K4 viral MIP protein.

SEQ ID NO 26 shows the cDNA nucleotide sequence of RRV ORF 16, a Bcl-2 homolog, corresponding to nucleotides 26846-27409 of SEQ ID NO 1, which has some similarity to KSHV ORF 16.

SEQ ID NO 27 shows the amino acid sequence of the RRV ORF 16 protein, a Bcl-2 protein homolog, which has some similarity to KSHV ORF 16 protein.

SEQ ID NO 28 shows the cDNA nucleotide sequence of RRV ORF 17, corresponding to the complement of nucleotides 27515-29125 of SEQ ID NO 1, encoding a capsid protein, which has some similarity to KSHV ORF 17.

SEQ ID NO 29 shows the amino acid sequence of the RRV ORF 17 protein, a capsid protein, which has some similarity to KSHV ORF 17 protein.

SEQ ID NO 30 shows the cDNA nucleotide sequence of RRV ORF 18, corresponding to nucleotides 28998-29897 of SEQ ID NO 1, which has some similarity to KSHV ORF 18.

SEQ ID NO 31 shows the amino acid sequence of the RRV ORF 18 protein, which has some similarity to KSHV ORF 18 protein.

SEQ ID NO 32 shows the cDNA nucleotide sequence of RRV ORF 19, corresponding to the complement of nucleotides 29905-31548 of SEQ ID NO 1, which encodes a tegument protein, and which has some similarity to KSHV ORF 19.

SEQ ID NO 33 shows the amino acid sequence of the RRV ORF 19 protein, a tegument protein, which has some similarity to KSHV ORF 19 protein.

SEQ ID NO 34 shows the cDNA nucleotide sequence of RRV ORF 20, corresponding to the complement of nucleotides 31043-32095 of SEQ ID NO 1, and which has some similarity to KSHV ORF 20.

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SEQ ID NO 35 shows the amino acid sequence of the RRV ORF 19 protein, which has some similarity to KSHV ORF 19 protein.

SEQ ID NO 36 shows the cDNA nucleotide sequence of RRV ORF 21, corresponding to nucleotides 32094-33767 of SEQ ID NO 1, which encodes a thymidine kinase protein, and which has some similarity to KSHV ORF 21.

SEQ ID NO 37 shows the amino acid sequence of the RRV ORF 21 protein, a thymidine kinase protein, which has some similarity to KSHV ORF 21 protein.

SEQ ID NO 38 shows the cDNA nucleotide sequence of RRV ORF 22, corresponding to nucleotides 33754-35868 of SEQ ID NO 1, and which encodes a glycoprotein H protein, and which has some similarity to KSHV ORF 22.

SEQ ID NO 39 shows the amino acid sequence of the RRV ORF 22 protein, a glycoprotein H protein, which has some similarity to KSHV ORF 22 protein.

SEQ ID NO 40 shows the cDNA nucleotide sequence of RRV ORF 23, corresponding to the complement of nucleotides 35865-37073 of SEQ ID NO 1, which has some similarity to KSHV ORF 23.

SEQ ID NO 41 shows the amino acid sequence of the RRV ORF 23 protein, which has some similarity to KSHV ORF 23 protein.

SEQ ID NO 42 shows the cDNA nucleotide sequence of RRV ORF 24, corresponding to the complement of nucleotides 37123-39321 of SEQ ID NO 1, and which has some similarity to KSHV ORF 24.

SEQ ID NO 43 shows the amino acid sequence of the RRV ORF 24 protein, which has some similarity to KSHV ORF 24 protein.

SEQ ID NO 44 shows the cDNA nucleotide sequence of RRV ORF 25, corresponding to nucleotides 39323-43459 of SEQ ID NO 1, which encodes a major capsid protein, and which has some similarity to KSHV ORF 25.

SEQ ID NO 45 shows the amino acid sequence of the RRV ORF 25 protein, a major capsid protein, which has some similarity to KSHV ORF 25 protein.

SEQ ID NO 46 shows the cDNA nucleotide sequence of RRV ORF 26, corresponding to nucleotides 43491-44408 of SEQ ID NO 1, which encodes a capsid protein, and which has some similarity to KSHV ORF 26.

SEQ ID NO 47 shows the amino acid sequence of the RRV ORF 26 protein, a capsid protein, which has some similarity to KSHV ORF 26 protein.

SEQ ID NO 48 shows the cDNA nucleotide sequence of RRV ORF 27, corresponding to nucleotides 44433-45242 of SEQ ID NO 1, and which has some similarity to KSHV ORF 27.

SEQ ID NO 49 shows the amino acid sequence of the RRV ORF 27 protein, which has some similarity to KSHV ORF 27 protein.

SEQ ID NO 50 shows the cDNA nucleotide sequence of RRV ORF 28, corresponding to

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nucleotides 45408-45683 of SEQ ID NO 1, and which has some similarity to KSHV ORF 28.

SEQ ID NO 51 shows the amino acid sequence of the RRV ORF 28 protein, which has some simil $\frac{1}{2}$ to KSHV ORF 28 protein.

SEQ ID NO 52 shows the cDNA nucleotide sequence of RRV ORF 29b, corresponding to the complement of nucleotides 45733-46779 of SEQ ID NO 1, and which has some similarity to KSHV ORF 29b.

SEQ ID NO 53 shows the amino acid sequence of the RRV ORF 29b, which has some similarity to KSHV ORF 29b protein.

SEQ ID NO 54 shows the cDNA nucleotide sequence of RRV ORF 30, corresponding to nucleotides 46905-47135 of SEQ ID NO 1, and which has some similarity to KSHV ORF 30.

SEQ ID NO 55 shows the amino acid sequence of the RRV ORF 30 protein, which has some similarity to KSHV ORF 30 protein.

SEQ ID NO 56 shows the cDNA nucleotide sequence of RRV ORF 31, corresponding to nucleotides 47093-47746 of SEQ ID NO 1, and which has some similarity to KSHV ORF 31.

SEQ ID NO 57 shows the amino acid sequence of the RRV ORF 31, protein which has some similarity to KSHV ORF 31 protein.

SEQ ID NO 58 shows the cDNA nucleotide sequence of RRV ORF 32, corresponding to nucleotides 47683-49077 of SEQ ID NO 1, and which has some similarity to KSHV ORF 32.

SEQ ID NO 59 shows the amino acid sequence of the RRV ORF 32 protein, which has some similarity to KSHV ORF 32 protein.

SEQ ID NO 60 shows the cDNA nucleotide sequence of RRV ORF 33, corresponding to nucleotides 49049-50059 of SEQ ID NO 1, and which has some similarity to KSHV ORF 33.

SEQ ID NO 61 shows the amino acid sequence of the RRV ORF 33 protein, which has some similarity to KSHV ORF 33 protein.

SEQ ID NO 62 shows the cDNA nucleotide sequence of RRV ORF 29a, corresponding to the complement of nucleotides 49977-50960 of SEQ ID NO 1, and which has some similarity to KSHV ORF 29a.

SEQ ID NO 63 shows the amino acid sequence of the RRV ORF 29a protein, which has some similarity to KSHV ORF 29a protein.

SEQ ID NO 64 shows the cDNA nucleotide sequence of RRV ORF 34, corresponding to nucleotides 50959-51942 of SEQ ID NO 1, and which has some similarity to KSHV ORF 34.

SEQ ID NO 65 shows the amino acid sequence of the RRV ORF 34 protein, which has some similarity to KSHV ORF 34 protein.

SEQ ID NO 66 shows the cDNA nucleotide sequence of RRV ORF 35, corresponding to nucleotides 51923-52372 of SEQ ID NO 1, and which has some similarity to KSHV ORF 35.

SEQ ID NO 67 shows the amino acid sequence of the RRV ORF 35 protein, which has some similarity to KSHV ORF 35 protein.

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SEQ ID NO 68 shows the cDNA nucleotide sequence of RRV ORF 36, corresponding to nucleotides 52278-53585 of SEQ ID NO 1, which encodes a kinase, and which has some similarity to KSHV ORF 36.

SEQ ID NO 69 shows the amino acid sequence of the RRV ORF 36 protein, a kinase, which has some similarity to KSHV ORF 36 protein.

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SEQ ID NO 70 shows the cDNA nucleotide sequence of RRV ORF 37, corresponding to nucleotides 53566-55008 of SEQ ID NO 1, which encodes an alkaline exonuclease, and which has some similarity to KSHV ORF 37.

SEQ ID NO 71 shows the amino acid sequence of the RRV ORF 37 protein, an alkaline exonuclease protein, which has some similarity to KSHV ORF 37 protein.

SEQ ID NO 72 shows the cDNA nucleotide sequence of RRV ORF 38, corresponding to nucleotides 54963-55172 of SEQ ID NO 1, and which has some similarity to KSHV ORF 38.

SEQ ID NO 73 shows the amino acid sequence of the RRV ORF 38 protein, which has some similarity to KSHV ORF 38 protein.

SEQ ID NO 74 shows the cDNA nucleotide sequence of RRV ORF 39, corresponding to the complement of nucleotides 55255-56391 of SEQ ID NO 1, which encodes glycoprotein M, and which has some similarity to KSHV ORF 39.

SEQ ID NO 75 shows the amino acid sequence of the RRV ORF 39 protein, glycoprotein M, which has some similarity to KSHV ORF 39 protein.

SEQ ID NO 76 shows the cDNA nucleotide sequence of RRV ORF 40, corresponding to nucleotides 56526-57932 of SEQ ID NO 1, which encodes helicase/primase, and which has some similarity to KSHV ORF 40.

SEQ ID NO 77 shows the amino acid sequence of the RRV ORF 40 protein, helicase/primase, which has some similarity to KSHV ORF 40 protein.

SEQ ID NO 78 shows the cDNA nucleotide sequence of RRV ORF 41, corresponding to nucleotides 57917-58528 of SEQ ID NO 1, which encodes helicase/primase, and which has some similarity to KSHV ORF 41.

SEQ ID NO 79 shows the amino acid sequence of the RRV ORF 41 protein, helicase/primase, which has some similarity to KSHV ORF 41 protein.

SEQ ID NO 80 shows the cDNA nucleotide sequence of RRV ORF 42, corresponding to the complement of nucleotides 58525-59343 of SEQ ID NO 1, which has some similarity to KSHV ORF 42.

SEQ ID NO 81 shows the amino acid sequence of the RRV ORF 42 protein, which has some similarity to KSHV ORF 42 protein.

SEQ ID NO 82 shows the cDNA nucleotide sequence of RRV ORF 43, corresponding to the complement of nucleotides 59297-61027 of SEQ ID NO 1, which encodes a capsid protein, and which has some similarity to KSHV ORF 43.

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SEQ ID NO 83 shows the amino acid sequence of the RRV ORF 43 protein, a capsid protein, which has some similarity to KSHV ORF 43 protein.

SEQ ID NO 84 shows the cDNA nucleotide sequence of RRV ORF 44, corresponding to nucleotides 60966-63338 of SEQ ID NO 1, which encodes helicase/primase, and which has some similarity to KSHV ORF 44.

SEQ ID NO 85 shows the amino acid sequence of the RRV ORF 44 protein, helicase/primase, which has some similarity to KSHV ORF 44 protein.

SEQ ID NO 86 shows the cDNA nucleotide sequence of RRV ORF 45, corresponding to the complement of nucleotides 63379-64437 of SEQ ID NO 1, and which has some similarity to KSHV ORF 45.

SEQ ID NO 87 shows the amino acid sequence of the RRV ORF 45 protein, which has some similarity to KSHV ORF 45 protein.

SEQ ID NO 88 shows the cDNA nucleotide sequence of RRV ORF 46, corresponding to the complement of nucleotides 64479-65246 of SEQ ID NO 1, which encodes uracil DNA glucosidase, and which has some similarity to KSHV ORF 46.

SEQ ID NO 89 shows the amino acid sequence of the RRV ORF 46 protein, uracil DNA glucosidase protein, which has some similarity to KSHV ORF 46 protein.

SEQ ID NO 90 shows the cDNA nucleotide sequence of RRV ORF 47, corresponding to the complement of nucleotides 65222-65731 of SEQ ID NO 1, which encodes glycoprotein L, which has some similarity to KSHV ORF 47.

SEQ ID NO 91 shows the amino acid sequence of the RRV ORF 47 protein, glycoprotein L, which has some similarity to KSHV ORF 47 protein.

SEQ ID NO 92 shows the cDNA nucleotide sequence of RRV ORF 48, corresponding to the complement of nucleotides 65999-67168 of SEQ ID NO 1, and which has some similarity to KSHV ORF 48.

SEQ ID NO 93 shows the amino acid sequence of the RRV ORF 48 protein, which has some similarity to KSHV ORF 48 protein.

SEQ ID NO 94 shows the cDNA nucleotide sequence of RRV ORF 49, corresponding to the complement of nucleotides 67398-68303 of SEQ ID NO 1, and which has some similarity to KSHV ORF 49.

SEQ ID NO 95 shows the amino acid sequence of the RRV ORF 49 protein, which has some similarity to KSHV ORF 49 protein.

SEQ ID NO 96 shows the cDNA nucleotide sequence of RRV ORF 50, corresponding to nucleotides 68494-70038 of SEQ ID NO 1, which encodes a transactivator, and which has some similarity to KSHV ORF 50.

SEQ ID NO 97 shows the amino acid sequence of the RRV ORF 50 protein, a transactivator protein, which has some similarity to KSHV ORF 50 protein.

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SEQ ID NO 98 shows the cDNA nucleotide sequence of RRV R4, corresponding to nucleotides 70355-70888 of SEQ ID NO 1.

SEQ ID NO 99 shows the amino acid sequence of the RRV R4 protein.

SEQ ID NO 100 shows the cDNA nucleotide sequence of RRV R5, corresponding to nucleotides 71468-72160 of SEQ ID NO 1.

SEQ ID NO 101 shows the amino acid sequence of the RRV R5 protein.

SEQ ID NO 102 shows the cDNA nucleotide sequence of RRV ORF 52, corresponding to the complement of nucleotides 72401-72820 of SEQ ID NO 1, and which has some similarity to KSHV ORF 52.

SEQ ID NO 103 shows the amino acid sequence of the RRV ORF 52 protein, which has some similarity to KSHV ORF 52 protein.

SEQ ID NO 104 shows the cDNA nucleotide sequence of RRV ORF 53, corresponding to the complement of nucleotides 72884-73198 of SEQ ID NO 1, and which has some similarity to KSHV ORF 53.

SEQ ID NO 105 shows the amino acid sequence of the RRV ORF 53 protein, which has some similarity to KSHV ORF 53 protein.

SEQ ID NO 106 shows the cDNA nucleotide sequence of RRV ORF 54, corresponding to nucleotides 73274-74146 of SEQ ID NO 1, which encodes a dUTPase, and which has some similarity to KSHV ORF 54.

SEQ ID NO 107 shows the amino acid sequence of the RRV ORF 54 protein, a dUTPase protein, which has some similarity to KSHV ORF 54 protein.

SEQ ID NO 108 shows the cDNA nucleotide sequence of RRV ORF 55, corresponding to the complement of nucleotides 74207-74839 of SEQ ID NO 1, and which has some similarity to KSHV ORF 55.

SEQ ID NO 109 shows the amino acid sequence of the RRV ORF 55 protein, which has some similarity to KSHV ORF 55 protein.

SEQ ID NO 110 shows the cDNA nucleotide sequence of RRV ORF 56, corresponding to nucleotides 74851-77337 of SEQ ID NO 1, which encodes a DNA replication protein, and which has some similarity to KSHV ORF 56.

SEQ ID NO 111 shows the amino acid sequence of the RRV ORF 56 protein, a DNA replication protein, which has some similarity to KSHV ORF 56 protein.

SEQ ID NO 112 shows the cDNA nucleotide sequence of RRV ORF 57, corresponding to nucleotides 77578-78906 of SEQ ID NO 1, which encodes an immediate-early gene product, and which has some similarity to KSHV ORF 57.

SEQ ID NO 113 shows the amino acid sequence of the RRV ORF 57 protein, a immediate-early gene product protein, which has some similarity to KSHV ORF 57 protein.

SEQ ID NO 114 shows the cDNA nucleotide sequence of RRV R6, corresponding to the

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complement of nucleotides 79266-80513 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEO ID NO 115 shows the amino acid sequence of the RRV R6 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 116 shows the cDNA nucleotide sequence of RRV R7, corresponding to the complement of nucleotides 80686-81933 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

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SEQ ID NO 117 shows the amino acid sequence of the RRV R7 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 118 shows the cDNA nucleotide sequence of RRV R8, corresponding to the complement of nucleotides 82262-83317 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEQ ID NO 119 shows the amino acid sequence of the RRV R8 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 120 shows the cDNA nucleotide sequence of RRV R9, corresponding to the complement of nucleotides 83491-84252 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEQ ID NO 121 shows the amino acid sequence of the RRV R9 protein, which has some similarity to KSHV vIRF K9 protein.

SEO ID NO 122 shows the cDNA nucleotide sequence of RRV R10, corresponding to the complement of nucleotides 85052-86209 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEO ID NO 123 shows the amino acid sequence of the RRV R10 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 124 shows the cDNA nucleotide sequence of RRV R11, corresponding to the complement of nucleotides 86355-87527 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEQ ID NO 125 shows the amino acid sequence of the RRV R11 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 126 shows the cDNA nucleotide sequence of RRV R12, corresponding to the complement of nucleotides 87894-88961 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

SEQ ID NO 127 shows the amino acid sequence of the RRV R12 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 128 shows the cDNA nucleotide sequence of RRV R13, corresponding to the complement of nucleotides 89122-90216 of SEQ ID NO 1, which has some similarity to KSHV vIRF K9 gene.

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SEQ ID NO 129 shows the amino acid sequence of the RRV R13 protein, which has some similarity to KSHV vIRF K9 protein.

SEQ ID NO 130 shows the cDNA nucleotide sequence of RRV ORF 58, corresponding to the complement of nucleotides 90462-91544 of SEQ ID NO 1, which has some similarity to KSHV ORF 58.

SEQ ID NO 131 shows the amino acid sequence of the RRV ORF 58 protein, which has some similarity to KSHV ORF 58 protein.

SEQ ID NO 132 shows the cDNA nucleotide sequence of RRV ORF 59, corresponding to the complement of nucleotides 91555-92739 of SEQ ID NO 1, which encodes a DNA replication protein, and which has some similarity to KSHV ORF 59.

SEQ ID NO 133 shows the amino acid sequence of the RRV ORF 59 protein, a DNA replication protein, which has some similarity to KSHV ORF 59 protein.

SEQ ID NO 134 shows the cDNA nucleotide sequence of RRV ORF 60, corresponding to the complement of nucleotides 92868-93812 of SEQ ID NO 1, which encodes a small ribonucleotide reductase protein, and which has some similarity to KSHV ORF 60.

SEQ ID NO 135 shows the amino acid sequence of the RRV ORF 60 protein, a small ribonucleotide reductase protein, which has some similarity to KSHV ORF 60 protein.

SEQ ID NO 136 shows the cDNA nucleotide sequence of RRV ORF 61, corresponding to the complement of nucleotides 93794-96160 of SEQ ID NO 1, which encodes a large ribonucleotide reductase protein, and which has some similarity to KSHV ORF 61.

SEQ ID NO 137 shows the amino acid sequence of the RRV ORF 61 protein, a large ribonucleotide reductase protein, which has some similarity to KSHV ORF 61 protein.

SEQ ID NO 138 shows the cDNA nucleotide sequence of RRV ORF 62, corresponding to the complement of nucleotides 96163-97158 of SEQ ID NO 1, which encodes a assembly/DNA maturation protein, and which has some similarity to KSHV ORF 62.

SEQ ID NO 139 shows the amino acid sequence of the RRV ORF 62 protein, a assembly/DNA maturation protein, which has some similarity to KSHV ORF 62 protein.

SEQ ID NO 140 shows the cDNA nucleotide sequence of RRV ORF 63, corresponding to nucleotides 97157-99976 of SEQ ID NO 1, which encodes a tegument protein, and which has some similarity to KSHV ORF 63.

SEQ ID NO 141 shows the amino acid sequence of the RRV ORF 63 protein, a tegument protein, which has some similarity to KSHV ORF 63 protein.

SEQ ID NO 142 shows the cDNA nucleotide sequence of RRV ORF 64, corresponding to nucleotides 99980-107626 of SEQ ID NO 1, which encodes a tegument protein, and which has some similarity to KSHV ORF 64.

SEQ ID NO 143 shows the amino acid sequence of the RRV ORF 64 protein, a tegument protein, which has some similarity to KSHV ORF 64 protein.

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SEQ ID NO 144 shows the cDNA nucleotide sequence of RRV ORF 65, corresponding to the complement of nucleotides 107637-108146 of SEQ ID NO 1, which encodes a capsid protein, and which has some similarity to KSHV ORF 65.

SEQ ID NO 145 shows the amino acid sequence of the RRV ORF 65 protein, a capsid protein, which has some similarity to KSHV ORF 65 protein.

SEQ ID NO 146 shows the cDNA nucleotide sequence of RRV ORF 66, corresponding to the complement of nucleotides 108152-109498 of SEQ ID NO 1, which has some similarity to KSHV ORF 66.

SEQ ID NO 147 shows the amino acid sequence of the RRV ORF 66 protein, which has some similarity to KSHV ORF 66 protein.

SEQ ID NO 148 shows the cDNA nucleotide sequence of RRV ORF 67, corresponding to the complement of nucleotides 109524-110198 of SEQ ID NO 1, which encodes a tegument protein, and which has some similarity to KSHV ORF 67.

SEQ ID NO 149 shows the amino acid sequence of the RRV ORF 67 protein, a tegument protein, which has some similarity to KSHV ORF 67 protein.

SEQ ID NO 150 shows the cDNA nucleotide sequence of RRV ORF 68, corresponding to nucleotides 110609-111982 of SEQ ID NO 1, which encodes a glycoprotein, and which has some similarity to KSHV ORF 68.

SEQ ID NO 151 shows the amino acid sequence of the RRV ORF 68 protein, a glycoprotein, which has some similarity to KSHV ORF 68 protein.

SEQ ID NO 152 shows the cDNA nucleotide sequence of RRV ORF 69, corresponding to nucleotides 112004-112897 of SEQ ID NO 1, which has some similarity to KSHV ORF 69.

SEQ ID NO 153 shows the amino acid sequence of the RRV ORF 69 protein, which has some similarity to KSHV ORF 69 protein.

SEQ ID NO 154 shows the cDNA nucleotide sequence of RRV ORF 71, corresponding to the complement of nucleotides 119211-119735 of SEQ ID NO 1, which encodes a FLIP protein, and which has some similarity to KSHV ORF 71.

SEQ ID NO 155 shows the amino acid sequence of the RRV ORF 71 protein, a FLIP protein, which has some similarity to KSHV ORF 71 protein.

SEQ ID NO 156 shows the cDNA nucleotide sequence of RRV ORF 72, corresponding to the complement of nucleotides 119794-120558 of SEQ ID NO 1, which encodes a cyclin D homolog, and which has some similarity to KSHV ORF 72.

SEQ ID NO 157 shows the amino acid sequence of the RRV ORF 72 protein, a cyclin D homolog protein, which has some similarity to KSHV ORF 72 protein.

SEQ ID NO 158 shows the cDNA nucleotide sequence of RRV ORF 73, corresponding to the complement of nucleotides 120866-122212 of SEQ ID NO 1, which encodes a latent nuclear antigen, and which has some similarity to KSHV ORF 73.

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SEQ ID NO 159 shows the amino acid sequence of the RRV ORF 73 protein, a latent nuclear antigen, which has some similarity to KSHV ORF 73 protein.

SEQ ID NO 160 shows the cDNA nucleotide sequence of RRV R15, corresponding to nucleotides 122866-123627 of SEQ ID NO 1, which has some similarity to KSHV K14 and ox-2.

SEQ ID NO 161 shows the amino acid sequence of the RRV R15 protein, which has some similarity to KSHV K14 and ox-2.

SEQ ID NO 162 shows the cDNA nucleotide sequence of RRV ORF 74, corresponding to nucleotides 123924-124952 of SEQ ID NO 1, which encodes a G protein coupled receptor, and which has some similarity to KSHV ORF 74.

SEQ ID NO 163 shows the amino acid sequence of the RRV ORF 74 protein, a G protein coupled receptor protein, which has some similarity to KSHV ORF 74 protein.

SEQ ID NO 164 shows the cDNA nucleotide sequence of RRV ORF 75, corresponding to the complement of nucleotides 125057-128953 of SEQ ID NO 1, which encodes a tegument protein, FGARAT, and which has some similarity to KSHV ORF 75.

SEQ ID NO 165 shows the amino acid sequence of the RRV ORF 75 protein, a tegument protein, FGARAT, which has some similarity to KSHV ORF 75 protein.

The cDNA sequences given in each of the even numbered sequences SEQ ID NOs 2-164 are the open reading frames of the RRV, with the nucleotide references in each of those sequences being given with reference to the nucleotide numbers set forth in SEQ ID NO 1.

SEQ ID NOs 166-172 are PCR primers used in the present invention.

SEQ ID NO 173 shows the coding sequence similar to that for MIP without AUG. Nucleic acid numbers correspond to those given in SEQ ID NO 1.

SEQ ID NOs 174-179 show the repeat regions of the RRV genome. Nucleic acid numbers correspond to those given in SEQ ID NO 1.

SEQ ID NOs 180-181 are probes specific for the KSHV thymidylate synthase (TS) gene used for Southern blot hybridization.

SEQ ID NOs 182-183 are oligonucleotide PCR primers specific for the RRV MIP gene.

ATCC DEPOSIT

A Budapest Treaty deposit of RRV 17577 was made with the American Type Culture Collection (ATCC), Manassas, Virginia, on March 12, 1998, and has been accorded ATCC Accession No. VR-2601.

DETAILED DESCRIPTION OF SEVERAL EMBODIMENTS

Abbreviations and Definitions

Animal: Living multicellular vertebrate organisms, a category which includes, for example, humans, non-human primates, mammals, and birds.

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Cell: A plant, animal, insect, bacterial, or fungal cell.

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Homologs: two nucleotide or amino acid sequences that share a common ancestral sequence and diverged when a species carrying that ancestral sequence split into two species. Homologs frequently show a substantial degree of sequence identity.

IL-6: Interleukin 6. IL-6 is a cytokine known to have pleiotropic immunological effects including anti-inflammatory and immunosuppressive effects (*Human Cytokines*, 1991, pg. 142-167, Blackwell Scientific Publications, Aggarwal and Gutterman, eds). Because IL-6 is a pleiotropic cytokine, IL-6 activity may be measured using a number of bioassays, including stimulation of immunoglobulin production in SKW6-CL4 cells as described by Hirano et al. (*Nature* 324:73-6, 1986) and stimulation of hybridoma cell growth as described by Matsuda et al., 1988 *Eur. J. Immunol.* 18:951-956, both of which are incorporated by reference. As used herein, the term "IL-6 biological activity" refers to the ability of a protein to show activity in at least one of these assay systems

Immuno-compromised: Lacking a normal immune response. Immuno-compromised refers to a condition in which some or all of an animal's immune system is inoperative, leaving the animal with an increased susceptibility to infection or disease. An animal may be rendered immuno-compromised by a biological agent such as, in the case of non-human primates, Simian Immunodeficiency Virus (SIV). Many strains of SIV have been isolated and characterized; any SIV strain that produces an immuno-compromised state can be used in the present invention including, but not limited to, for example, SIVmac239 (Kestler et al., 1990, Science 248: 1109-12). SIVmac251 (Daniels et al., 1985, Science 228: 1201-4), SIVdeltaB670 (Murphy-Corb et al., 1986, Nature 321:435) and SIVmne (Benveniste et al., 1988, J. Virol. 62:2091-101). In addition, hybrid SIV/HIV chimeras as known in the field can be employed, as can HIV-2. Simian type D retroviruses (SRVs) which cause an AIDS-like disease in rhesus monkeys, can alternatively be used to immuno-compromise the animals in place of SIV. These viral agents are administered to the animal using conventional means, such as intravenous or intramuscular injection, or oral, intrarectal or intravaginal inoculation (also see Example 24). Either intact viral particles or viral DNA may be administered. As known in the field, plasmid constructs containing the entire SIV genome are infectious when inoculated into animals and so may be employed in place of purified viral DNA.

Alternatively, an animal may be rendered immuno-compromised by administration of agents that target the immune system, including byt not limited to anti-CD3 antibody (CD3 being the T-cell receptor) either alone or conjugated with a toxic moiety, or immunosuppressive compounds including prednisone, azathioprine, cyclosporine A, and cyclophosphamide. Where an immunosuppressive compound such as cyclosporine is employed, an allogenic stimulus (such as a blood transfusion) may be administered with the subsequent administration of RRV to activate infection.

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Alternatively, other methods of rendering an animal immuno-compromised may be used, including radiation treatment and surgical intervention.

Isolated: An "isolated" biological component (such as a nucleic acid, peptide or protein) has been substantially separated, produced apart from, or purified away from other biological components in the cell of the organism in which the component naturally occurs, i.e., other chromosomal and extrachromosomal DNA and RNA, and proteins. Nucleic acids, peptides and proteins which have been "isolated" thus include nucleic acids and proteins purified by standard purification methods. The term also embraces nucleic acids, peptides and proteins prepared by recombinant expression in a host cell as well as chemically synthesized nucleic acids.

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KSHV: Kaposi's sarcoma-associated herpesvirus. KSHV is a herpesvirus associated with (and thought to be the etiological agent of) Kaposi's sarcoma in humans.

Lymphoproliferative Disorder: a group of disorders characterized by proliferation of lymphoid tissue, such as lymphocytic leukemia and malignant lymphoma, and characterized by such signs as lymphocytosis, lymphadenopathy, and splenomegaly.

MIP: macrophage inflammatory protein. The acronym MIP is used to describe a family of cytokines that includes MIP1 (Davatelis et al., 1989, Science 243: 1066-8) and MIP2 (U.S. Patent No. 5,145,676). MIPs mediate pleiotropic immunological effects including activation of neutrophils to undergo an oxidative burst. MIPs are also intrinsically pyrogenic. MIP biological activity can be detected and quantified using bioassays as described in Kedal et al. (Science 277:1656-9, 1997) and Boshoff et al. (Science 278:290-4, 1997) that measure MIP concentrations using HIV inhibition and calcium mobilization, respectively. As used herein, the term "MIP biological activity" refers to the ability of a protein to show activity in at least one of these assay systems.

Non-human primate: Simian primates including chimpanzees, orangutans, baboons, and macaques. Any non-human primate may be used to produce a KSHV-disease animal model by the methods disclosed herein. Thus, in addition to the rhesus macaque models described in detail below, pigtail and cynomologus macaques and baboons may also be used to produce KSHV-disease animal models by the methods disclosed herein.

Oligonucleotide: A linear polynucleotide sequence of up to about 200 nucleotide bases in length, for example a polynucleotide (such as DNA or RNA) which is at least 6 nucleotides, for example at least 15, 25, 50, 100 or even 200 nucleotides long.

Operably linked: A first nucleic acid sequence is operably linked with a second nucleic acid sequence when the first nucleic acid sequence is placed in a functional relationship with the second nucleic acid sequence. For instance, a promoter is operably linked to a coding sequence if the promoter affects the transcription or expression of the coding sequence. Generally, operably linked DNA sequences are contiguous and, where necessary to join two protein coding regions, in the same reading frame.

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ORF: open reading frame. Contains a series of nucleotide triplets (codons) coding for amino acids without any termination codons. These sequences are usually translatable into protein.

PCR: polymerase chain reaction. Describes a technique in which cycles of denaturation, annealing with primer, and then extension with DNA polymerase are used to amplify the number of copies of a target DNA sequence.

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Pharmaceutically acceptable carriers: The pharmaceutically acceptable carriers useful in this invention include conventional carriers. Remington's Pharmaceutical Sciences, by E. W. Martin, Mack Publishing Co., Easton, PA, 15th Edition (1975), describes compositions and formulations suitable for pharmaceutical delivery of the viruses, nucleic acids and/or proteins herein disclosed.

In general, the nature of the carrier will depend on the particular mode of administration being employed. For instance, parenteral formulations usually comprise injectable fluids that include pharmaceutically and physiologically acceptable fluids such as water, physiological saline, balanced salt solutions, aqueous dextrose, glycerol, ethanol, combinations thereof, or the like, as a vehicle. The carrier and composition can be sterile, and the formulation suits the mode of administration. For solid compositions (e.g., powder, pill, tablet, or capsule forms), conventional non-toxic solid carriers can include, for example, pharmaceutical grades of mannitol, lactose, starch, sodium saccharine, cellulose, magnesium carbonate, or magnesium stearate. In addition to biologically-neutral carriers, pharmaceutical compositions to be administered can contain minor amounts of non-toxic auxiliary substances, such as wetting or emulsifying agents, preservatives, and pH buffering agents and the like, for example sodium acetate or sorbitan monolaurate.

The composition can be a liquid solution, suspension, emulsion, tablet, pill, capsule, sustained release formulation, or powder. The composition can be formulated as a suppository, with traditional binders and carriers such as triglycerides.

Probes and primers: Nucleic acid probes and primers may readily be prepared based on the amino acid sequences provided by this invention. A probe is an isolated nucleic acid attached to a detectable label or reporter molecule. Typical labels include radioactive isotopes, ligands, chemiluminescent agents, and enzymes. Methods for labeling and guidance in the choice of labels appropriate for various purposes are discussed, e.g., in Sambrook et al., in Molecular Cloning: A Laboratory Manual, Cold Spring (1989) and Ausubel et al., in Current Protocols in Molecular Biology, Greene Publishing Associates and Wiley-Intersciences (1987).

Primers are short nucleic acids, such as DNA oligonucleotides 10 nucleotides or more in length. Primers may be annealed to a complementary target DNA strand by nucleic acid hybridization to form a hybrid between the primer and the target DNA strand, and then extended along the target DNA strand by a DNA polymerase enzyme. Primer pairs can be used for amplification of a nucleic acid sequence, e.g., by the polymerase chain reaction (PCR) or other

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nucleic-acid amplification methods known in the art.

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Methods for preparing and using probes and primers are described, for example, in Sambrook et al. (Molecular Cloning: A Laboratory Manual, 2nd ed., vol. 1-3, Cold Spring Harbor, New York. 1989); Ausubel et al. (Current Protocols in Molecular Biology, Greene Publ. Assoc. & Wiley-Intersciences. 1987) and Innis et al. (PCR Protocols, A Guide to Methods and Applications, 1990, Innis et al. (eds.), 21-27, Academic Press, Inc., San Diego, California). PCR primer pairs can be derived from a known sequence, for example, by using computer programs intended for that purpose such as Primer (Version 0.5, © 1991, Whitehead Institute for Biomedical Research, Cambridge, MA).

Probes and primers as used in the present invention typically comprise at least 15 contiguous nucleotides of the RRV genome sequence (SEQ ID NO 1). One of skill in the art will appreciate that the specificity of a particular probe or primer increases with its length. Thus, for example, a primer comprising 20 consecutive nucleotides will anneal to a target with a higher specificity than a corresponding primer of only 15 nucleotides. Thus, in order to obtain greater specificity, probes and primers may be selected that comprise 20, 25, 30, 35, 40, 50 or more consecutive nucleotides. In order to enhance specificity, longer probes and primers may also be employed, such as probes and primers that comprise at least 20, 30, 40, 50, 60, 70, 80, 90, 100, or 150 consecutive nucleotides of the disclosed nucleic acid sequences.

Alternatively, such probes and primers may comprise at least 15, 20, 30, 40, 50, 60, 70, 80, 90, 100, or 150 consecutive nucleotides that share a defined level of sequence identity with the disclosed RRV sequence, for instance, at least a 50%, 60%, 70%, 80%, 90%, 95% or 98% sequence identity. Alternatively, such probes and primers may be nucleotide molecules which hybridize under wash conditions of 70°C and about 0.2 x SSC for 1 hour, or alternatively under less stringent conditions of 65°C, 60°C, or 55°C with from about 0.2 to 2 x SSC (with, for instance, about 0.1% SDS) for 1 hour with a portion of the RRV sequence.

Purified: The term purified does not require absolute purity; rather, it is intended as a relative term. Thus, for example, a purified peptide preparation is one in which the peptide or protein is more enriched than the peptide or protein is in its natural environment within a cell. Preferably, a preparation is purified such that the protein or peptide represents at least 50% of the total peptide or protein content of the preparation.

Recombinant: A recombinant nucleic acid is one that has a sequence that is not naturally occurring or has a sequence that is made by an artificial combination of two otherwise separated segments of sequence. This artificial combination is often accomplished by chemical synthesis or, more commonly, by the artificial manipulation of isolated segments of nucleic acids, e.g., by genetic engineering techniques.

RRV 17577: Rhesus macaque rhadinovirus RRV isolate 17577. A Budapest Treaty deposit of RRV 17577 was made with the American Type Culture Collection, Manassas, Virginia,

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on March 12, 1998, and has been accorded ATCC Accession No. VR-2601. This virus may be grown on primary rhesus fibroblasts, as described below (see Examples 1 and 14), using standard virological techniques. Alternatively, it may be grown on commercially available rhesus cell lines, including those available from ATCC, such as ATCC CRL-6306 and ATCC CL-160. Infection of a non-human primate with RRV 17577 may be accomplished using any standard method, including intravenous injection (see Examples 13, 23 and 24). Typically, infection is achieved by intravenous injection of around 10⁶ plaque forming units (PFUs) of RRV 17577.

RRV: A virus having the virological and immunological characteristics of RRV 17577, and which causes Kaposi's sarcoma in immunocompromised Rheusus monkeys which are infected with the virus. In particular examples, the RRV has at least 85% (for example at least 90%, 95% or 98%) sequence identity to SEQ ID NO 1.

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Sequence Identity: The similarity between two nucleic acid sequences, or two amino acid sequences, is expressed in terms of the similarity between the sequences, otherwise referred to as sequence identity. Sequence identity is frequently measured in terms of percentage identity (or similarity or homology); the higher the percentage, the more similar the two sequences are. Homologs or orthologs of the RRV proteins and the corresponding DNA sequences, will possess a relatively high degree of sequence identity when aligned using standard methods. This homology will be more significant when the orthologous proteins or DNAs are derived from species which are more closely related (e.g., human and chimpanzee sequences), compared to species more distantly related (e.g., human and C. elegans sequences).

Typically, RRV orthologs are at least 50% identical at the nucleotide level and at least 50% identical at the amino acid level when comparing RRV to an orthologous RRV sequences.

Methods of alignment of sequences for comparison are well known in the art. Various programs and alignment algorithms are described in: Smith & Waterman, Adv. Appl. Math. 2:482, 1981; Needleman & Wunsch, J. Mol. Biol. 48:443, 1970; Pearson & Lipman, Proc. Natl. Acad. Sci. USA 85:2444, 1988; Higgins & Sharp, Gene, 73:237-44, 1988; Higgins & Sharp, CABIOS 5:151-3, 1989; Corpet et al., Nuc. Acids Res. 16:10881-90, 1988; Huang et al. Computer Appls. in the Biosciences 8, 155-65, 1992; and Pearson et al., Meth. Mol. Bio. 24:307-31, 1994. Altschul et al. (Nature Genetics 6:119-29, 1994), presents a detailed consideration of sequence alignment methods and homology calculations.

The NCBI Basic Local Alignment Search Tool (BLAST) (Altschul et al. *J. Mol. Biol.* 215:403-10, 1990) is available from several sources, including the National Center for Biotechnology Information (NCBI, Bethesda, MD) and on the Internet, for use in connection with the sequence analysis programs blastp, blastn, blastx, tblastn and tblastx. It can be accessed at http://www.ncbi.nlm.nih.gov/BLAST/. A description of how to determine sequence identity using this program is available at http://www.ncbi.nlm.nih.gov/BLAST/blast_help.html. As used herein, sequence identity is commonly determined with the BLAST software set to default

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parameters. For instance, blastn (version 2.0.6) software may be used to determine sequence identity between two nucleic acid sequences using default parameters. For comparison of two polypeptides, blastp (version 2.0.6) software may be used with default parameters.

An alternative alignment tool is the ALIGN Global Optimal Alignment tool (version 3.0) available from Biology Workbench at http://biology.ncsa.uiuc.edu. This tool may be used with settings set to default parameters to align two known sequences. References for this tool include Meyers and Miller (*CABIOS* 4:11-7, 1989).

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Homologs of the disclosed RRV nucleic acids typically possess at least 50% sequence identity counted over the length of one of the nucleic acids (the reference nucleic acid) using the NCBI Blast 2.0.6, gapped blastn set to default parameters. Nucleic acids showing substantial similarity when assessed by this method may show, for example, at least 50%, 60%, 70%, 80%, 90%, 95% or even 98% or greater sequence identity. When less than the entire sequence is being compared for sequence identity, substantially similar nucleotide sequences will typically possess at least 70% sequence identity over short windows of 30-90 nucleic acids, and may possess sequence identities of at least 80%, 90%, 95% or 98% or greater.

Homologs of the disclosed RRV proteins typically possess at least 50% sequence identity counted over full-length alignment with the amino acid sequence of RRV using the NCBI Blast 2.0, gapped blastp set to default parameters. For comparisons of amino acid sequences of greater than about 30 amino acids, the Blast 2 sequences function is employed using the default BLOSUM62 matrix set to default parameters, (gap existence cost of 11, and a per residue gap cost of 1). When aligning short peptides (fewer than around 30 amino acids), the alignment should be performed using the Blast 2 sequences function, employing the PAM30 matrix set to default parameters (open gap 9, extension gap 1 penalties). Proteins with even greater similarity to the reference sequence will show increasing percentage identities when assessed by this method, such as at least 50%, at least 55%, at least 60%, at least 70%, at least 75%, at least 80%, at least 90%, at least 95%, at least 98%, or at least 99% sequence identity. When less than the entire sequence is being compared for sequence identity, homologs will typically possess at least 70% sequence identity over short windows of 10-20 amino acids, and may possess sequence identities of at least 75%, at least 85% or at least 90%, at least 95% or 98% depending on their similarity to the reference sequence. Methods for determining sequence identity over such short windows are described at http://www.ncbi.nlm.nih.gov/BLAST/blast FAQs.html.

When comparing degrees of sequence identity between similar proteins, the degree of identity will be equal to or less than that the degree of similarity, due to the fact the similarity takes into account conservative amino acid substitutions. So, for instance, the degree of sequence identity between to substantially similar proteins may be at least 43%, 50%, 55%, 65%, 75%, 85%, 95%, 98% or more.

One of ordinary skill in the art will appreciate that these sequence identity ranges are

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provided for guidance only; it is entirely possible that strongly significant homologs could be obtained that fall outside of the ranges provided. The present invention provides not only the peptide hereologs that are described above, but also nucleic acid molecules that encode such homologs.

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An alternative indication that two nucleic acid molecules are closely related is that the two molecules hybridize to each other under stringent conditions, as described in Example 23.

Specific binding agent: An agent that binds substantially only to a defined target. As used herein, the term "RRV peptide specific binding agent" includes anti-RRV peptide antibodies and other agents that bind substantially only to the RRV peptide. Such "peptide specific binding agents" include anti-IL-6 and anti-MIP antibodies. The antibodies may be monoclonal or polyclonal antibodies that are specific for an RRV peptide, as well as immunologically effective portions ("fragments") thereof. In one embodiment, the antibodies used in the present invention are monoclonal antibodies (or immunologically effective portions thereof) and may also be humanized monoclonal antibodies (or immunologically effective portions thereof).

Immunologically effective portions of monoclonal antibodies include Fab, Fab', F(ab')₂, Fabc and Fv portions (for a review, see Better and Horowitz, *Methods. Enzymol.* 178:476-96, 1989). Anti-inhibitory peptide antibodies may also be produced using standard procedures described in a number of texts, including <u>Antibodies, A Laboratory Manual</u> by Harlow and Lane, Cold Spring Harbor Laboratory (1988).

Methods of making humanized monoclonal antibodies are well known, and include those described in U.S. Patent Nos. 5,585,089; 5,565,332; 5,225,539; 5,693,761; 5,693,762; 5,585,089; and 5,530,101 and references cited therein. Similarly, methods of making and using immunologically effective portions of monoclonal antibodies, also referred to as antibody fragments, are well known and include those described in Better and Horowitz, 1989, *Meth. Enzymol.* 178:176-496; Better et al., 1990, Better and Horowitz, 1990, Advances in Gene technology: The Molecular Biology of Immune Disease & the Immune Response (ICSU Short Reports); Glockshuber et al., 1990, *Biochemistry* 29:1362-7; and U.S. Patent Nos. 5,648,237; 4,946,778 and 5,455,030, and references cited therein.

The determination that a particular agent binds substantially only to an RRV peptide may readily be made by using or adapting routine procedures. One suitable *in vitro* assay makes use of the Western blotting procedure (described in many standard texts, including Antibodies, A Laboratory Manual by Harlow and Lane). Western blotting may be used to determine that a given RRV peptide binding agent, such as an anti-IL-6 or MIP peptide monoclonal antibody, binds substantially only to the specific RRV protein.

Subject: This term includes both human and non-human subjects. Similarly, the term "patient" includes both human and veterinary subjects.

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Transformed: A transformed cell is a cell into which has been introduced a nucleic acid molecule by molecular biology techniques. As used herein, the term transformation encompasses all techniques by which a nucleic acid molecule might be introduced into such a cell, including transfection with viral vectors, transformation with plasmid vectors, and introduction of naked DNA by electroporation, lipofection, and particle gun acceleration.

Transgenic Cell: Transformed cells which contain foreign, non-native DNA.

Variants of Amino Acid and Nucleic Acid Sequences: The production of RRV proteins can be accomplished in a variety of ways (for example see Examples 17, 21 and 25). DNA sequences which encode the protein, or a fragment of the protein, can be engineered such that they allow the protein to be expressed in eukaryotic cells, bacteria, insects, and/or plants. In order to accomplish this expression, the DNA sequence can be altered and operably linked to other regulatory sequences. The final product, which contains the regulatory sequences and the therapeutic protein, is referred to as a vector. This vector can then be introduced into the eukaryotic cells, bacteria, insect, and/or plant. Once inside the cell the vector allows the protein to be produced.

One of ordinary skill in the art will appreciate that the DNA can be altered in numerous ways without affecting the biological activity of the encoded protein. For example, PCR may be used to produce variations in the DNA sequence which encodes RRV proteins, such as IL-6 or MIP. Such variants may be variants that are optimized for codon preference in a host cell that is to be used to express the protein, or other sequence changes that facilitate expression.

Two types of cDNA sequence variant may be produced. In the first type, the variation in the cDNA sequence is not manifested as a change in the amino acid sequence of the encoded polypeptide. These silent variations are simply a reflection of the degeneracy of the genetic code. In the second type, the cDNA sequence variation does result in a change in the amino acid sequence of the encoded protein. In such cases, the variant cDNA sequence produces a variant polypeptide sequence. In order to preserve the functional and immunologic identity of the encoded polypeptide, it is preferred that any such amino acid substitutions are conservative. Conservative substitutions replace one amino acid with another amino acid that has some homology in size, hydrophobicity, etc. Such substitutions generally are conservative when it is desired to finely modulate the characteristics of the protein. For example, conservative substitutions generally maintain (a) the structure of the polypeptide backbone in the area of the substitution, for example, as a sheet or helical conformation, (b) the charge or hydrophobicity of the molecule at the target site, or (c) the bulk of the side chain.

Examples of amino acids which may be substituted for an original amino acid in a protein and which are regarded as conservative substitutions include: Ser for Ala; Lys for Arg; Gln or His for Asn; Glu for Asp; Ser for Cys; Asn for Gln; Asp for Glu; Pro for Gly; Asn or Gln for His; Leu or Val for Ile; Ile or Val for Leu; Arg or Gln for Lys; Leu or Ile for Met; Met, Leu or Tyr

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for Phe; Thr for Ser; Ser for Thr; Tyr for Trp; Trp or Phe for Tyr; and Ile or Leu for Val.

The substitutions which in general are expected to produce the greatest changes in protein properties will be non-conservative, for instance changes in which (a) a hydrophilic residue, e.g., seryl or threonyl, is substituted for (or by) a hydrophobic residue, e.g., leucyl, isoleucyl, phenylalanyl, valyl or alanyl; (b) a cysteine or proline is substituted for (or by) any other residue; (c) a residue having an electropositive side chain, e.g., lysyl, arginyl, or histadyl, is substituted for (or by) an electronegative residue, e.g., glutamyl or aspartyl; or (d) a residue having a bulky side chain, e.g., phenylalanine, is substituted for (or by) one not having a side chain, e.g., glycine.

Variations in the DNA sequence that result in amino acid changes, whether conservative or not, should be minimized in order to preserve the functional and immunologic identity of the encoded protein. The immunologic identity of the protein may be assessed by determining whether it is recognized by an antibody to an RRV protein; a variant that is recognized by such an antibody is immunologically conserved. Any DNA sequence variant will preferably introduce no more than 20, and preferably fewer than 10 amino acid substitutions into the encoded polypeptide. Variant amino acid sequences can, for example, be 80%, 90%, 95% or even 98% identical to the native amino acid sequence.

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Vector: A nucleic acid molecule as introduced into a host cell, thereby producing a transformed host cell. A vector may include nucleic acid sequences that permit it to replicate in the host cell, such as an origin of replication. A vector may also include one or more selectable marker genes and other genetic elements known in the art.

Virion: A complete viral particle including envelope, capsid (if any), and nucleic acid elements.

The present invention utilizes standard laboratory practices for the cloning, manipulation and sequencing of nucleic acids, purification and analysis of proteins and other molecular biological and biochemical techniques, unless otherwise stipulated. Such techniques are explained in detail in standard laboratory manuals such as Sambrook et al. (Molecular Cloning: A Laboratory Manual, 2nd ed., vol. 1-3, Cold Spring Harbor, New York. 1989) and Ausubel et al. (Current Protocols in Molecular Biology, Greene Publ. Assoc. & Wiley-Intersciences. 1987).

EXAMPLE 1

Isolation of RRV

This example describes how RRV was isolated from a rhesus macaque monkey. Fresh, dispersed bone marrow (BM) cells were isopynic gradient-purified (Ficoll-Paque, Pharmacia) from a 2 yr, 202 day old captive-reared rhesus macaque that was euthanized 503 days after intravenous infection with an SIVmac239 variant. Gradient-purified BM mononuclear cells were seeded into T-25 culture flasks and grown in the presence of Endothelial SFM media (GIBCO) supplemented

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with 10% fetal bovine serum, 1% L-glutamine, 1% penicillin-streptomycin-neomycin and 30 μ g/mL endothelial cell growth supplement.

Cultures developing cytopathic effects (CPE) were rapidly frozen in liquid N₂ and thawed, and supernatants clarified by centrifugation and filtered through a 0.45 μ membrane. Filtered extracts were then used as inoculum on primary rhesus macaque fibroblast cultures. Fibroblast cultures developing CPE were scraped free into medium, pelleted at 400 xg, washed in phosphate-buffered saline and suspended in cold Ito and Karnovsky's fixative (2.5% glutaraldehyde, 0.5% picric acid, 1.6% paraformaldehyde, 0.005% ruthenium red) in 0.1 M sodium cacodylate buffer, pH 7.4 for 2 hours. Fixed cells were washed in cacodylate buffer, post-fixed in 1% OsO₄ and 0.8% K₃Fe (Cn)₆ in cacodylate buffer for 1 hour, rinsed in distilled H₂O and pre-stained in 2% aqueous uranyl acetate for 1 hour. Fixed and pre-stained cells were dehydrated in a graded series of acetone imbedded in Epon 812 epoxy resin, polymerized at 60°C and sectioned at 60 nm on an MT 5000 ultramicrotome. Copper grid mounted sections were stained with lead citrate and Uranyl acetate and viewed on a Phillips 300 electron microscope.

By electron microscopy, numerous herpesvirus particles were observed in the cells. This macaque developed LPD characterized as lymphocytic masses in myeloid and nonlymphoid tissues which were confirmed histopathologically as solid pleomorphic lymphoid masses.

EXAMPLE 2

20 Initial Characterization of RRV

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Infectious virus was purified from infected primary rhesus fibroblast cultures exhibiting 100% CPE (see Example 1). Infected cells were harvested and disrupted by freeze-thawing and the cell debris removed by low speed centrifugation. Supernatants were centrifuged in a Beckman JA-14 rotor for 1 hour at 9000 rpm to pellet the virus, which was further purified through a six-step sorbitol gradient ranging from 20 to 70%, spun in a Beckman SW41 rotor for 2 hours at 18,000 rpm. Virus was diluted in balanced buffered salts solution and then pelleted through a 20% sorbitol cushion. Pelleted virus was resuspended in Tris-EDTA buffer (TE; 10 mM Tris-HCl, pH 8.0, and 1 mM EDTA) and lysed in TE with 0.6% SDS and proteinase K (200 μ g) at 37°C for 5 hours. Viral DNA was then isolated by CsCl2 gradient centrifugation in a Beckman Ti 75 rotor at 38.4 K rpm for 72 hours, collected and dialyzed against TE.

The viral DNA was analyzed using degenerate primer polymerase chain reaction (PCR) amplification and Southern blot hybridization with a probe specific for the KSHV thymidylate synthase (TS) gene: (5'-CTATACTGCCATTTC-3', SEQ ID NO 180 and 5'-ATGTTTCCGTTTGTA-3', SEQ ID NO 181). The probe itself had the sequence of the KSHV TS (Orf 70 gene). Four genes were identified by these methods. A fragment encoding a portion of the viral DNA polymerase was obtained and DNA sequence analysis revealed that the virus was most likely a gamma herpesvirus, as amino acid sequence identity was highest among this class of

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herpesviruses. DNA sequence analysis of the viral DNA fragment found to hybridize to the KSHV TS probe revealed three open reading frames (ORFs) with homology to KSHV (Nicholas et al., 1997, *Nature Med.* 3:287-92; Russo et al.; 1996, *Proc Natl Acad Sci USA* 93:14862-7). One ORF encodes a homologue of macrophage inflammatory protein MIP-1 with amino acid sequence identity with KSHV MIP-II, the second ORF encodes a thymidylate synthase homologue and the third ORF encodes a homologue of interleukin-6 (IL-6) with homology to the rhesus IL-6 and KSHV IL-6. The presence of an IL-6-like cytokine and an MIP-1-like CC-chemokine flanking TS resembles the genomic organization of KSHV, indicating this virus is related to KSHV and is referred to herein as rhesus rhadinovirus (RRV).

To determine if RRV is present in tissue containing the lymphocytic masses, oligonucleotide PCR primers specific for the RRV MIP gene (vMIP-1, 5'-CCTATGGGCTCCATGAGC-3', SEQ ID NO 182; and vMIP-2, 5'-ATCGTCAATCAGGCTGCG-3', SEQ ID NO 183) were designed in an attempt to detect viral DNA in tissue from the macaque. By semi-quantitative PCR analysis, viral DNA sequences were detected in DNA samples from bone marrow at approximately 590 copies per 0.1 µg of tissue DNA. Because rhesus macaques held in captivity are reported to be naturally infected with a herpesvirus similar to KSHV, bone marrow DNA samples were isolated from normal and SIVmac239-infected macaques without LPD and analyzed by PCR. There was no evidence of viral DNA sequences. Additionally, since simian Epstein-Barr virus (EBV) has been found to be present in high copy number in lymphomas from SIV-infected macaques (Baskin et al., 1986, *J. Natl. Cancer Inst.* 77:127-39; Feichtinger et al., 1990, *Amer. J. Pathol.* 137:1311-5), the tissue samples from the macaque with disease were also analyzed by PCR for rhesus EBV (RhEBV) using oligonucleotide primers for RhEBV latent membrane protein 1. By this analysis, no signal for RhEBV was detected, suggesting that the RRV may be a contributing factor for LPD in this SIV-infected macaque.

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EXAMPLE 3

Preparation of RRV DNA for Cloning

Primary rhesus fibroblasts grown in two 850 cm 2 roller bottles were infected with RRV at an MOI of 0.1 and the virus was harvested from the culture supernatant and the infected monolayers 10 to 12 days post-infection. Cellular debris was removed from the culture supernatant by centrifugation at 1,000 x g for 10 minutes. Intracellular virus particles were released by sonication followed by centrifugation to pellet debris.

The two clarified supernatants were then combined and the virus was pelleted by centrifugation at 12,500 x g for 1 hour at 4°C, and further purified through a six-step sorbitol gradient ranging from 20 to 70%. Gradients were centrifuged in a Beckman SW41 rotor for 2 hours at 18,000 rpm at 4°C. The interface containing the virus was collected and diluted with cold buffered saline solution. The virus was then pelleted by centrifugation in the SW41 for 50

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minutes at 18,000 rpm. The virus pellet was resuspended in 9.2 ml of TE (see Example 2) before the addition of 0.6 ml of 10% sodium dodecylsulfate (SDS) and 0.2 ml of proteinase K (10 mg/mL) to release the viral DNA. Viral DNA was isolated by CsCl₂ gradient centrifugation in a Beckman Ti75 rotor at 38,400 rpm for 72 hours, collected, and dialyzed against TE.

To ensure that the DNA isolated contained all the necessary sequences required for RRV replication, DNA was transfected, in duplicate, into primary rhesus fibroblasts by the calcium phosphate method without dimethyl sulfoxide shock and observed for cytopathic effects (CPE). Control transfections, lacking viral DNA or calcium phosphate, did not develop CPE.

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EXAMPLE 4

Construction of the Cosmid Library

Approximately $100 \mu g$ of purified RRV DNA (Example 3) was partially digested with Sau3A I. Aliquots taken at various time points were run on a 0.5% agarose gel and examined for the fraction which gave the desired range of fragments (30 - 42 kb). The selected fraction was dephosphorylated with calf intestinal alkaline phosphatase and $1 \mu g$ ligated into the cosmid vector SuperCos 1, prepared essentially as described by the manufacturer (Stratagene, La Jolla, CA). The resulting ligation product was packaged using GigaPack II Gold packaging extract (Stratagene) and grown for the isolation of recombinant cosmids.

Individual recombinant cosmids were grown in 3 ml cultures and the cosmid DNA was isolated by alkaline lysis. Cosmid DNA was digested with EcoR1 and the DNA fragments separated on a 0.8% agarose gel. The separated fragments were transferred to nitrocellulose and probed with various PCR amplification products corresponding to specific KSHV ORFs. Hybridization of the probes to the transferred recombinant cosmids was done under conditions of moderate stringency (2x SSC-0.1%SDS at 55°C) with each of the KSHV-specific probes and at high stringency (0.2x SSC-0.1%SDS at 60°C) with the RRV-specific probes. By this analysis and restriction endonuclease mapping, the recombinant cosmids were aligned and a set of recombinants was identified that represented the entire viral genome when compared to digested viral DNA.

EXAMPLE 5

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Cloning and Sequencing

Ten micrograms of each purified recombinant cosmid (Example 4) were digested with EcoRI and the resulting fragments isolated from a 0.8% agarose gel using the QiaQuick gel extraction protocol (Qiagen). Recovered fragments were ligated into pSP73 (Promega). Individual clones were selected by restriction enzyme screening of DNA recovered by alkaline lysis from overnight cultures. Sequencing templates were prepared by alkaline lysis, followed by precipitation with 6.5% polyethylene glycol and 0.8 M NaCl. Templates were resuspended at a concentration of $0.1 \mu g/\mu l$ and end sequences were determined using primers corresponding to the

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SP6 and T7 promoters of pSP73. Internal sequences were determined using a combination of subcloning using convenient restriction sites and custom primers. DNA sequencing reactions were performed with the Applied Biosystems (ABI) PRISM Dye Terminator Cycle Sequencing Ready Reaction kits with AmpliTaq DNA polymerase per the manufacturer's instructions. Sequence data was acquired using an ABI 373A Sequencer in the Molecular Biology Core at the Oregon Regional Primate Research Center. The primary EcoRI fragments were sequentially arranged by sequencing across the EcoRI sites in the intact cosmids using custom primers. Except for those regions containing long, high GC repeat units, the entire viral DNA sequence was determined with a redundancy of 3- to 4-fold.

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Sequences not accessible to custom primers or restriction subcloning were determined following deletion subcloning using the Exo Size Deletion kit (New England Biolabs). To accommodate this protocol, fragments were subcloned into vectors with restriction sites capable of generating the needed 3' and 5' overhanging ends. Double restriction digests to generate 3' and 5' overhanging ends were performed on 10 µg of recombinant plasmid DNA, which was then subjected to exonuclease III digestion. Aliquots were removed from the exonuclease III digests at empirically-determined time points, frozen on dry ice, then, after all the time points had been collected, incubated for 15 minutes at 65°C to inactivate the enzyme. The DNA was then treated with Mung bean nuclease (MBN) for 30 minutes at 30°C. Prior to addition of 3 µl of MBN to the 12 µl exonuclease III product, the enzyme was diluted 1/25 to reduce nonspecific digestion.

Nuclease-treated DNA was recovered using the Wizard prep system (Promega), then incubated for 30 minutes with 2.5 units of T4 DNA polymerase (Life Technologies) and 1 µM dNTPs at 37°C. The final product was ligated overnight with T4 DNA ligase and used to transform competent XL1 blue bacteria. Deletion products were size selected by restriction digests of DNA recovered from 3 ml cultures.

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EXAMPLE 6

Assembly of the RRV Sequence, Assignment of ORFs, and Nomenclature

Factura (ABI) and Autoassembler (ABI) were used to assemble the final sequence from individual sequencing runs. Open reading frames in the RRV sequence were determined with the program MacVector (Oxford Molecular Group), using a setting of 100 or more amino acids. Putative ORFs were then translated and compared to a database of KSHV ORFs. RRV ORFs which matched KSHV ORFs were then compared to GenBank using BLASTP to verify the similarity, followed by a Gap analysis (Wisconsin GCG analysis package; Oxford Molecular Group) to determine the levels of similarity and identity between the RRV and KSHV proteins. When a gap in the genome of RRV corresponded to the location of a KSHV ORF with less than 100 amino acids, MacVector was reset to a lower limit. RRV ORFs were assigned the names of HVS ORFs when they showed similarity to KSHV ORFs with the same name.

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The nucleotide sequence data from this study have been deposited in the GenBank, EMBL, and DDBJ nucleotide sequence databases under accession number AF083501 (SEQ ID NO 1).

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EXAMPLE 7

Primary Structure Of the RRV Genome

The genomic nucleotide sequence of the RRV genome (as shown in SEQ ID NO 1) was determined using twenty-nine EcoRI fragments (as shown in FIG. 2) from seven overlapping isolates of a partial Sau3A I cosmid library. Cosmids were selected by hybridization with PCR products from KSHV ORFs. EcoRI fragments from each cosmid were subcloned into pSP73 (Promega) and sequenced. The EcoRI fragments were arranged in the proper order by sequencing across the EcoRI junctions in the parent cosmids using custom primers. Greater than 98% of the viral genome was determined on both strands. The average sequencing redundancy was between 3 and 4, but three regions were sequenced on only one strand. One of these regions is a 106 bp segment of ORF 61 (SEQ ID Nos 136 and 137) that was blocked on one side by an apparent hairpin. This segment was sequenced multiple times in one direction using templates derived from independent overlapping cosmids. The other two regions are 1 kb, high G + C, repetitive sequences. These segments, which are discussed in more detail below, were sequenced completely on one strand using a combination of custom primers and exonuclease III deletions.

Terminal repeats were identified on both the left and right ends of the genome and the sequence between them was designated as the LUR of the genome. The first base to the right of the left terminal repeat was designated base one. The LUR is 133,719 bp long (SEQ ID NO 1). The G+C content of RRV is 52.2%, which is comparable to the 53.5% G+C content of KSHV, but considerably higher than the 34.5% G+C content of the HVS genome. The CpG ratio is 1.11, which is substantially higher than the ratio found for other gamma-herpesviruses.

ORFs were identified by MacVector and compared to a database containing the full complement of known KSHV ORFs. Matches between RRV and KSHV proteins were verified by a BLASTP search of GenBank with the RRV proteins and then by Gap analysis. The initial screening for ORFs used a minimum size limit of 100 amino acids. This limit was reduced when smaller KSHV ORFs existed in locations corresponding to unassigned regions of RRV. Using this approach, 82 ORFs were identified (even-numbered SEQ ID Nos 2-164), with 67 of these corresponding to ORFs found in both KSHV and HVS. In accordance with the standard nomenclature for rhadinoviruses, these ORFs were labeled according to the HVS designation. The 15 ORFs not found in HVS were assigned labels beginning with R (for rhesus), indicating their presence in RRV, but not HVS. Some of these genes have counterparts in KSHV.

A map of the genome of RRV is presented in FIG. 3, with all identified ORFs and their orientations. The BamHI, EcoRI, and HindIII restriction sites in relation to the genome are shown

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in FIG. 2. The BamHI and Hind III maps were generated from the final compiled sequence. The EcoRI map was also generated from the final compiled sequence, but it was further characterized by sequencing across the EcoRI junctions in the parent cosmids. Fragment sizes for each restriction map are presented in FIG. 4.

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EXAMPLE 8

Genomic Organization of RRV

The overall genomic organization of RRV matches the general structure of gamma-herpesviruses, with blocks of shared ORFs interrupted at specific locations (referred to as divergent loci) where the viral genomes code for acquired cellular genes. The primate rhadinoviruses form a subset of the gamma-herpesviruses and their genomes are correspondingly more similar to each other than to other members of the family.

The genomic sequence of RRV is presented in SEQ ID NO 1. FIG. 3 shows a schematic representation of the ORFs of RRV with a corresponding restriction map. FIG. 4 shows the location, size and description of the RRV ORFs.

EXAMPLE 9

Comparison of RRV and KSHV ORFs

A comparison of corresponding repeats in RRV and KSHV is shown in FIG. 5. In addition, FIG. 5 presents data for RRV ORFs along with the results of the Gap analysis of ORFs shared by RRV, KSHV, and HVS. All HVS-like ORFs found in KSHV are found in RRV. A comparison table of interferon regulatory elements encoded by the RRV and KSHV genomes is shown in FIG. 6.

EXAMPLE 10

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Comparison of RRV and HVS ORFs

FIG. 7 shows the results of the Gap analysis of ORFs shared by RRV, KSHV, and HVS. In general, RRV and HVS ORFs are highly similar when the corresponding RRV and KSHV ORFs are highly similar, although the Gap values are generally lower.

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EXAMPLE 11

ORFs Unique to RRV and KSHV

RRV includes 14 genes which are not found in HVS (R1 SEQ ID NOS 2 and 3; R2 SEQ ID NOS 20 and 21; R3 SEQ ID NOS 24 and 25; R4 SEQ ID NOS 98 and 99; R5 SEQ ID NOS 100 and 101; R6 SEQ ID NOS 114 and 115; R7 SEQ ID NOS 116 and 117; R8 SEQ ID NOS 118 and 119; R9 SEQ ID NOS 120 and 121; R10 SEQ ID NOS 122 and 123; R11 SEQ ID NOS 124 and 125; R12 SEQ ID NOS 126 and 127; R13 SEQ ID NOS 128 and 129; and R15 SEQ ID NOS 160 and 161). These are designated in FIG. 3 as "R" ORFs. Of these fifteen genes, 11 have

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counterparts in the genome of KSHV. R2 (SEQ ID NOs 20 and 21) and R3 (SEQ ID NOs 24 and 25) are cytokine genes. R2 has functional homology to K2, the vIL-6 gene of KSHV. Gap analysis of the vIL-6 genes from KSHV and RRV shows no notable similarity, but both possess four conserved cysteines found in cellular IL-6. In addition, RRV vIL-6 has IL-6-like activity in cell culture. R3 has a small, but clear, similarity to KSHV K4, a vMIP1β gene. It is the only vMIP gene in RRV, as compared to the three vMIP genes found in KSHV.

RRV R6 through R13 are vIRFs as are KSHV K9 through K11 (FIG. 6). K9, the most studied of the KSHV vIRFs, does not have a DNA binding domain, but has been demonstrated to inhibit the endogenous cellular interferon response pathways. Five of the RRV vIRFs (R6, R7, R8, R10, and R11) are similar to K9, though only R10 has a similarity greater than 30%. The remaining similarities fall between 26% and 30%. There is no measurable similarity between any RRV vIRF and any KSHV vIRF other than K9. There is, however, a pattern of higher similarity between members of the RRV vIRF family. R6, R7, R8, and R9 are most similar to R10, R11, R12, and R13, respectively, with the similarities falling between 50% and 62%. The pattern of similarity suggests a single, possibly recent, gene duplication event for RRV which increased the number of vIRFs in the genome from four to eight.

The final RRV gene with a unique KSHV counterpart is R15, which has some similarity to K14, a viral NCAM Ox-2 homologue. The similarity between R15 and K14 (35.2%) is relatively low compared to most other shared proteins.

A number of genes in RRV appear to be truly unique. R1 colocalizes with, but has no similarity to, K1, a KSHV gene that has been demonstrated to have *in vivo* transforming ability. K1 and R1 both colocalize with ORF1, or STP (saimiri transforming protein), although both K1 and R1 are in opposite orientations compared to STP. A BLASTP search of GenBank using R1 reveals a limited amino-terminal similarity to a series of Fc receptors, including a potential transmembrane domain. These data suggest that R1, like K1 and STP, may have transforming potential via transmembrane signaling.

R4 and R5 are located between ORF 50 and 52, the same location as K8 and K8.1 in KSHV; however, there is no similarity between either R4 or R5 and the KSHV proteins. A BLASTP search of GenBank failed to show any significant alignments with either R4 or R5, so their functions are unknown.

RRV has no confirmed ORFs in the region corresponding to K12, the ubiquitously expressed kaposin gene. A large ORF exists to the right of ORF 71, but it has no apparent control regions (TATA box or polyadenylation signal), so it has not been designated as a true ORF, pending identification of transcripts from this region. No ORFs corresponding to KSHV K15 have been identified.

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EXAMPLE 12

Co-localization of Repeat Units in RRV and KSHV

The RRV genome contains three highly repetitive regions, which correspond to three of the repetitive regions of KSHV: frnk, zppa, and mdsk (FIG. 5). KSHV frnk and zppa, and the corresponding RRV repetitive regions, rDL-B and rDL-E, respectively, are tandem repeats.

The first element of the RRV syko repeat is much lower in G + C content than the corresponding KSHV element, although the sizes are comparable (FIG. 5). The second element is over 700 bp longer than the corresponding KSHV element. The first element of the RRV vrigo repeat is 30% longer than the corresponding KSHV element, and the second RRV element is over four times as long as the second KSHV element. There is no sequence similarity between the various elements of the two viruses nor is there any similarity between any two repeat sequences in RRV.

Not all repeat elements found in KSHV have corresponding repeats in RRV. This includes the KSHV vnct and waka/jwka repeats. This also includes the moi repeat, which is located in the center of the KSHV ORF 73 and is responsible for the divergent lengths of RRV and KSHV ORF 73. Moi is described in the annotations to the KSHV GenBank entry as having 15 different 11-16 bp repeats. The result of this repeat element is the presence in ORF 73 of a highly acidic central domain, with a large number of glutamate residues coded by a repeating GAG codon. KSHV ORF 73 is a potential leucine zipper protein, with a number of leucine zipper sites in the repeat region. RRV lacks the moi repeat and its concomitant acidic domain. It also lacks any evidence for a leucine zipper, indicating that the biology of ORF 73 in RRV may be substantially different than the biology of ORF 73 in KSHV.

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Production of Simian Kaposi's Sarcoma (KS) and Lymphoproliferative Disorders Model

This example describes how the RRV cloned above can be used to produce a non-human primate model for Kaposi's sarcoma and lymphoproliferative disorders. Four rhesus macaques (identification numbers 18483, 18503, 18540 and 18570) that were approximately 1.5 years old, and PCR- and seronegative for RRV were selected. To perform the antibody analysis, infected cells were solubilized with 0.5% Nonidet P-40 and 1% sodium deoxycholate in phosphate buffered saline, and clarified in a Beckman SW28 rotor at 23,500 rpm for 1 hour at 4°C. The clarified supernatant was used as antigen for coating enzyme-linked immunosorbent assay (ELISA) plates (500 ng/well). ELISAs were then performed essentially as described by Kodama et al. (AIDS Res Hum Retroviruses 5:337-43, 1989).

All of the animals were then inoculated intravenously with cell-free supernatants containing the equivalent of 5 ng of p27 prepared from COS-1 cells transfected with an

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SIVmac239 molecular clone (Endres et al., 1995, SW. J Med. Primatol. 24:141-4). The PBMCs from all macaques were prescreened for in vitro susceptibility to virus infection as described by Naidu et al. (J. Virol. 62:4691-6, 1988). All inoculations and animal manipulations were performed according to institutional guidelines at the Oregon Regional Primate Research Center (Beaverton, OR). Every 3-4 days for 4 weeks, then at 2-week intervals, macaques were sedated with ketamine hydrochloride (10 mg per kilogram of body weight) and examined for fever, weight loss, cutaneous signs, lymphadenopathy, and hepatomegaly or splenomegaly. At these times, venipuncture was performed and blood specimens collected. Plasma was monitored for virus during the first 4 weeks with the SIV p27 enzyme-linked immunosorbent assay (ELISA) (Coulter Corp. Hialeah, FL.). T cell subsets and B cells were measured by flow cytometry with the OKT4 (CD4, Ortho), B9.11 (CD8, Coulter), and B-Ly-1 (CD20, Coulter) monoclonal antibodies.

At 8 weeks post-SIV infection, rhesus macaques 18483 and 18570 were inoculated intravenously with 5 x 106 plaque forming units of gradient purified RRV that was grown and titered by plaque assay on primary rhesus fibroblasts. The two remaining macaques (18503 and 18540) were kept as SIV-infected controls. Every 3-4 days for 2 weeks, once a week for 4 weeks, then at 2 week intervals, the macaques were examined and blood samples collected and analyzed. Virus isolations were performed by cocultivation of 2 x 10⁵ PBMCs from each of the macaques with primary rhesus fibroblasts in duplicate. Cell cultures were monitored every 2-3 days for 3-4 weeks for cytopathic effects characteristic of RRV. PBLs were also analyzed by PCR for the presence of viral DNA. PCR analysis for RRV was performed with the following oligonucleotide primers: vMIP-1, 5' CCTATGGGCTCCATGAGC 3' (SEQ ID NO 166); and vMIP-2, 5' ATCGTCAATCAGGCTGCG 3' (SEQ ID NO 167). The conditions for PCR were 94°C for 2 minutes (1 cycle); 94°C for 0.5 minutes, 50°C for 0.5 min, 72°C for 0.5 minutes (30 cycles); 72°C extension for 5 minutes (1 cycle). Each PCR reaction used 0.1 Fg of total DNA, 50 pmole of each primer, 1 U of Vent polymerase, 40 µM each of deoxynucleotide triphosphate, 10 mM KCl, 10 mM Tris-HCl (pH 8.8), 10 mM (NH₄)₂SO₄, 2 mM MgSO₄ and 0.1% Triton X-100 in a final volume of 50 µL. The PCR reactions were run out on a 1% agarose gel, transferred to nitrocellulose, and probed with a 32P-ATP-labeled oligonucleotide primer specific for vMIP-3 (5' ATATTAAACACTCGCCGC- 3' SEQ ID NO 168). Hybridizations were performed overnight at room temperature in 6X SSC, 0.1% SDS and 10 µg/mL E. coli tRNA. Southern blots were then washed with 2X SSC and 0.1% SDS twice at room temperature followed by two washes for 1 hour in 2X SSC and 0.1% SDS at 47°C. Bound probe was visualized by exposing NEN duPont reflection film to the washed membrane at 80°C with an NEN duPont Reflection screen.

Infectious RRV was recovered from the peripheral blood mononuclear cells (PBMCs) of both RRV macaques injected with RRV as early as 4 weeks after inoculation for one macaque (18570) and 8 weeks for the other macaque (18483), but not from the control macaques. The peripheral blood leukocytes (PBL) from both macaques were also shown to harbor viral DNA as

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determined by PCR and Southern blot analysis for the viral MIP gene, as early as 4 weeks after inoculation for one macaque (18483) and as late as 14 weeks for the second macaque (18570). Additionally, antibody responses to RRV were observed as measured by ELISA in the RRV-infected macaques beginning 4 weeks post-infection, but not in the control macaques.

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Flow cytometry analysis (FACS) of PBLs at the indicated weeks post-infection (FIGS 8A-8D) showed there was limited CD4+ lymphocyte depletion after SIV infection in both groups of macaques followed by a rebound and sustained CD4+ lymphocyte counts. However, examination of CD20+ B lymphocytes revealed significant differences between the two groups. The two control macaques exhibited a dramatic and sustained decline in CD20+ B lymphocytes (FIGs. 8C and 8D), whereas both co-infected macaques exhibited a transient increase in B lymphocytes beginning 6 weeks after RRV infection (FIGs. 8A and 8B). The increase in CD20+ B lymphocytes correlated with the isolation and/or detection of RRV in both macaques; however, viral load did not appear to correlate with the increase in CD20+ B lymphocytes when all samples from each macaque were analyzed simultaneously. It has been reported that CD23, a B cell activation marker, is induced by RhEBV infection of macaques (Moghaddam et al. 1997, Science 276:2020-33). FACS analysis of PBMCs from RRV-infected macaques revealed no detectable CD23+ cells. This would suggest that the mechanism responsible for increased numbers of CD20+ B lymphocytes following RRV infection differ from the activation of B lymphocytes by RhEBV.

Routine physical examinations were performed on all four macaques, and early symptoms of SIV infection were observed in all four macaques by 2 weeks, including fever, rash and malaise. However, 11 weeks after inoculation with RRV, macaques 18483 and 18570 developed marked lymphadenopathy and splenomegaly, estimated to be enlarged 10 to 20 times the size of a normal spleen. In contrast, there was only slight lymph node enlargement in the control macaques not infected with RRV and no detectable enlargement of the spleen. Lymph node biopsies of the RRV-infected macaques revealed almost identical histology, characterized by a predominately follicular lesion with giant germinal centers and paracortical hyperplasia with increased vascularity, resembling angiofollicular lymph node hyperplasia associated with KSHV in Castleman's disease (Lachant et al. 1985, Am. J. Clin. Pathol. 83:27-33). In contrast, the lymph nodes of the control macaques exhibited atrophied lymphoid follicles and paracortical depletion characteristic of SIV-induced lymphoid atrophy (Chalifoux et al., 1987, Am. J. Pathol. 128:104-10; Ringler et al., 1989, Am. J. Pathol. 134:373-83; Wyand et al, 1989, Am. J. Pathol. 134:385-93). By FACS analysis, the majority of the lymph node mononuclear cells were CD20+ B lymphocytes in RRV-infected macaques, whereas CD4+ and CD8+ T lymphocytes predominated in the control macaques.

The presence of viral DNA was determined by PCR analysis on DNA derived from PBLs.

Detection of antibodies to RRV was determined by enzyme-linked immunosorbent assay (ELISA)

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on plates coated with extracts derived from RRV-infected cells. By PCR analysis, RRV sequences were more prevalent in the lymph nodes than in the peripheral blood of RRV-infected macaques, whereas control macaques were negative for RRV sequences (FIGs. 9A and 9B).

Additional disease manifestations were also observed in the RRV-infected macaques that parallel clinical features and B cell abnormalities observed in AIDS patients. Hypergammaglobulinemia was observed in the RRV-infected macaque that the virus was derived from, as well as in the macaques experimentally infected with RRV, whereas the two control macaques had gammaglobulin levels similar to those before SIV infection. In addition, one of two RRV-infected macaques (18570) developed severe autoimmune hemolytic anemia 30 weeks after RRV infection, a condition frequently observed in MCD patients (Parravicini et al., 1997, Am. J. Pathol. 151:1517-22).

The second of the two RRV-infected macaques developed other unique clinical manifestations that paralleled those of AIDS patients with KS. At 60 weeks post-RRV infection it developed a distended abdomen that was clinically evident upon physical examination. Palpation revealed a pronounced fluid accumulation in the peritoneal cavity. This animal was euthanized due to persistent fluid accumulation and hyperbilirubinemia. Necropsy analysis on this animal revealed an abundance of ascites fluid, which was comprised predominately of CD20 B cells, as identified by FACS analysis. In addition, this animal exhibited a mesenchymal proliferative lesion throughout the viscera, that was identified by histopathological examination to be retroperitoneal fibromatosis (RF). RF is an abnormal highly vascularized mesenchymal proliferative lesion that exhibits histological features resembling Kaposi's Sarcoma. Analysis of DNA isolated from the ascites and RF lesion by PCR with RRV MIP primers (given in Example 2) revealed a high viral load, implying RRV infection was responsible for these abnormal proliferations.

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EXAMPLE 14

Other Methods to Prepare RRV Nucleic Acid Sequences

Obtaining the RRV Viral Genome

The RRV genome of the invention (SEQ ID NO 1) can be procured by *de novo* isolation from a viral culture. A biological sample of the virus (accession number VR-2601) may be obtained from the ATCC in Manassas, VA. This virus can be grown *in vitro* using primary rhesus fibroblasts (see Example 1). The virus is harvested from the culture supernatant and the infected host cells. Cellular debris is removed by centrifugation and intracellular virus particles may be released by sonication followed by centrifugation to pellet debris. The virus is then pelleted by centrifugation and further purified through a six-step sorbitol gradient. The interface containing the virus is collected and the virus then pelleted by centrifugation, and the viral DNA released by SDS disruption. Viral DNA may be isolated by CsCl₂ gradient centrifugation.

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Obtaining Selected Polynucleotides from the Viral genome

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The isolated viral genome can be used as a source of polynucleotides as identified by the sequence as disclosed herein (SEQ ID NO 1). The polymerase chain reaction (PCR) may be used to amplify any polynucleotide selected from the known viral sequence using the viral genome as a source of template DNA. The template DNA may also be provided in the form of one or more cosmids that contain fragments of the viral genome. Alternately, cDNA, produced by reverse transcription of RNA extracted from RRV infected host cells, may be used as a template in a reverse-transcription PCR (RT-PCR) reaction. Methods and conditions for PCR and RT-PCR amplification are described in Innis et al. (PCR Protocols, A Guide to Methods and Applications, 1990, Innis et al. (eds.), 21-27, Academic Press, Inc., San Diego, California).

The selection of PCR primers may be made according to the portions of the genome to be amplified. Primers may be chosen to amplify small fragments of the genome, ORFs or fragments including many contiguous genes from the genome. Variations in amplification conditions may be required to accommodate primers of differing lengths, and such considerations are well known in the art and are discussed in Innis at al. (PCR Protocols, A Guide to Methods and Applications, 1990, Innis et al. (eds.), 21-27, Academic Press, Inc., San Diego, California), Sambrook et al. (Molecular Cloning: A Laboratory Manual, 2nd ed., vol. 1-3, Cold Spring Harbor, New York. 1989) and Ausubel et al. (Current Protocols in Molecular Biology, Greene Publ. Assoc. & Wiley-Intersciences. 1987). For example, the ORF corresponding to the MIP gene may be amplified from an RRV genomic (or appropriate cosmid) template using the following pair of primers: 5' ATGAGGGGCCTTTTCGTGTGC 3' (SEQ ID NO 169) and 5' CTGAATCCCGCTGCCAAGGCC 3' (SEQ ID NO 170).

Likewise, the ORF corresponding to the IL-6 gene may be amplified from an RRV genomic (or appropriate cosmid) template using the following pair of primers: 5'
ATGTTCCCTGTCTGGTTCGTC 3' (SEQ ID NO 171) and 5' TTACATCATAGCTATTGCGCG
3' (SEQ ID NO 172).

Such primers are illustrative only and it will be readily appreciated by one of ordinary skill in the art that many different primers may be selected from the sequence disclosed and used in PCR amplification reactions to amplify DNA sequences of interest from the RRV genome.

Polynucleotides that may be obtained by the above methods include, for example: the entire polynucleotide genome of RRV as shown in SEQ ID NO 1; ORFs of this genome; oligonucleotides comprising at least 15, 20, 30, 40, 50, 70, 100 and 150 consecutive nucleotides of the genome sequence as shown in SEQ ID NO 1; nucleic acid sequences defined by nucleotides 1 to 11031 of SEQ ID NO 1 and nucleotides 21625 to 133719 of SEQ ID NO 1; and ORFs selected from these nucleic acid sequences. It is readily apparent that fragments of any length may be made using the above methods and information.

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EXAMPLE 15

Therapeutic and Diagnostic Uses of the RRV IL-6 Protein

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As disclosed herein, the genome of RRV possesses an IL-6 gene (FIGS. 1 and 10 and SEQ ID NO 20) similar to that found in KSHV. The IL-6 and MIP proteins of KSHV are thought to be important in disease pathology, such as in Kaposi's sarcoma. The primary structure of the RRV IL-6 protein is shown in FIG. 10 (SEQ ID NO 21). Given this sequence information, one can readily make derivative proteins of RRV IL-6. Such derivative proteins include proteins that differ from the primary amino acid sequence as shown in FIG. 10 (SEQ ID NO 21) by one or more conservative amino acid substitutions. Examples of such conservative substitutions are given in the DEFINITIONS section of the specification. Derivative proteins also include proteins consisting of an amino acid sequence that has a defined degree of amino acid similarity with the RRV IL-6 protein. For instance, such derivative proteins will typically have at least 50% sequence similarity (and may have at least 60%, 70%, 80%, 90%, 95%, 98% or even 99% sequence similarity) with the RRV IL-6 protein. Such derivative proteins will not only share sequence similarity with KSHV IL-6 but will also possess IL-6 biological activity.

IL-6 is a cytokine known to have pleiotropic immunological effects including antiinflammatory and immunosuppressive effects, and may be used in several therapeutic and
diagnostic applications. RRV IL-6 of the invention may be likewise be used. For instance, IL-6
may be used to induce stimulation of hematopoietic stem cells, and to encourage proliferation,
differentiation and terminal maturation of erythroid cells from hematopoietic cells. Thus, for
instance, RRV IL-6 may be used *in vivo* or *ex vivo* to treat diseases that involve leukopenia and
thrombocytopenia. Such uses include stimulation of hematopoietic cells of radiotherapy patients or
people exposed to radiation accidentally. IL-6 may be used in such applications in conjunction
with GM-CSF (granulocyte-macrophage stimulating factor) (see U.S. Patent Nos. 5,610,056 and
5,620,685, herein incorporated by reference). IL-6 can also be used to stimulate growth of
megakaryocytes and platelets, and for the inhibition of tumor growth (see U.S. Patent No.
5,620,685, herein incorporated by reference). IL-6 can also be used for the treatment of
leukemias, such as chronic myeloid leukemia (CML) and acute myeloid leukemia, by inducing
terminal differentiation of cells with IL-6 (see WO 90/01943, herein incorporated by reference).
RRV IL-6 may be used for all such applications.

Therapeutic applications may involve the administration of RRV IL-6 in a number of ways. RRV IL-6 may be administered *in vivo*, e.g., by injection systemically or locally, for instance, into a subject. Many other forms of *in vivo* administration are possible including intravenous, subcutaneous, across a mucous membrane (anally, vaginally or sublingually), transdermal or by direct injection. Additionally, it may be administered *ex vivo*, by the removal of cells from a subject, the treatment of these cells *in vitro* with RRV IL-6, and the replacement of

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these cells into the subject. Another recently developed method of delivery of a protein drug is by introducing the gene coding for the drug into a subject, for instance within the genome of a virus, such as an adenovirus or a retrovirus, whereby the protein is expressed in the subject. Other modes of administration are provided in Example 25.

Such examples are provided for illustrative purposes only and it will be seen that RRV IL-6 may be used in a variety of topical and systemic immunological treatments where it would be desirable to stimulate cell proliferation or to induce anti-inflammatory or immunosuppressive effects. Additionally, IL-6 of the invention may be used for research and diagnostic purposes as discussed generally herein. For instance, IL-6 may be used to produce antibodies for diagnostic purposes to diagnose diseases characterized by increased or decreased production of IL-6, and the nucleic acid sequence encoding IL-6 may be used to produce probes and primers for diagnostic and research purposes or for gene therapy applications. The IL-6 could also be used as a targeting molecule for identifying cells with receptors for IL-6, and for directing therapeutic agents to these cells, for example by linking detector or therapeutic molecules to IL-6.

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EXAMPLE 16

Therapeutic and Diagnostic Uses of the RRV MIP Protein

The genome of RRV as disclosed herein possesses an MIP gene (FIGS. 1 and 11 and SEQ ID NO 24) similar to that found in KSHV. The primary structure of the RRV MIP protein is shown in FIG. 11 (SEQ ID NO 25). Given this sequence information, one can readily make derivative proteins of RRV MIP. Such derivative proteins include proteins that differ from the primary amino acid sequence as shown in FIG. 11 (SEQ ID NO 25) by one or more conservative amino acid substitutions. Derivative proteins also include proteins consisting of an amino acid sequence that has a defined degree of amino acid similarity with the RRV MIP protein. Typically such derivative proteins will have at least 50% sequence similarity with the RRV MIP protein, and may have at least 60%, 70%, 80%, 90%, 95%, 98%, or even 99% sequence similarity. Such derivative proteins will not only share sequence similarity with KSHV MIP but will also possess MIP biological activity. MIP biological activity can be detected and quantified using bioassays as described in Kedal et al. (Science 277:1656-9, 1997) and Boshoff et al. (Science 278:290-4, 1997) that measure MIP concentrations using HIV inhibition and calcium mobilization, respectively.

MIP is a cytokine that activates neutrophils to undergo an oxidative burst and is also intrinsically pyrogenic. The MIP genes and proteins of the invention may be used in several therapeutic and diagnostic ways. The RRV MIP protein may be used for the same applications as other MIP proteins. Treatment of wounds to promote healing by application of MIP to the wound site is discussed in U.S. Patent No. 5,145,676. U.S. Patent No. 5,474,983 (herein incorporated by reference) discusses various methods of treatment of inflammatory diseases including asthma,

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allergies and dermatitis. U.S. Patent No. 5,656,724 (herein incorporated by reference) discloses the use of MIP to suppress proliferation of dividing myeloid cells e.g., for the treatment of neutropenia. Use of MIP to inhibit HIV is discussed by Kedal et al. (*Science* 277:1656-9, 1997). RRV MIP may be used for all such applications.

As illustrated for IL-6 above, MIP may be administered in various ways to provide a therapeutic effect including *in vivo*, *ex vivo* and by gene therapy.

Such examples are provided for illustrative purposes only and it will be seen that MIP may be used in a variety of topical, systemic, in vivo and ex vivo immunological treatments where it would be desirable to activate neutrophils or to induce fever. Additionally, MIP of the invention may be used for diagnostic purposes as discussed generally herein. For instance, MIP may be used to produce antibodies for diagnostic purposes to diagnose diseases characterized by increased or decreased production of MIP, and the nucleic acid sequence encoding MIP may be used to produce probes and primers for diagnostic and research purposes, or for gene therapy applications.

The MIP could also be used as a targeting molecule for identifying cells with receptors for MIP, and for directing therapeutic agents to these cells, for example by linking detector or therapeutic molecules to MIP.

Although Examples 15 and 16 provide examples of therapeutic uses of the RRV IL-6 and MIP proteins, any of the other proteins encoded by the RRV can also be administered therapeutically, or diagnostically. For example, RRV proteins that induce pathological or physiological conditions in a recipient can be administered to stimulate that condition for study, or to provide an animal or human model of the condition. That model can then be used to study the condition, or treatments for it.

EXAMPLE 17

Expression of RRV cDNA Sequences

With the provision of the RRV genomic (SEQ ID NO 1) and cDNAs (even-numbered SEQ ID Nos 2-164), the expression and purification of any of the RRV proteins (odd-numbered SEQ ID Nos 3-165), from any species, by standard laboratory techniques is now enabled. Fragments amplified as described herein can be cloned into standard cloning vectors and expressed in commonly used expression systems consisting of a cloning vector and a cell system in which the vector is replicated and expressed. Purified proteins may be used for functional analyses, antibody production, diagnosis, and patient therapy. Furthermore, the DNA sequences of the RRV cDNAs can be manipulated in studies to understand the expression of RRV genes and the function of their products. Mutant forms of RRV may be isolated based upon information contained herein, and may be studied in order to detect alteration in expression patterns in terms of relative quantities, and functional properties of the encoded mutant RRV protein. Partial or full-length cDNA sequences, which encode for the protein, may be ligated into bacterial expression vectors. Methods for expressing large amounts of protein from a cloned gene introduced into E. coli may

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be utilized for the purification, localization and functional analysis of proteins. For example, fusion proteins consisting of amino terminal peptides encoded by a portion of the *E. coli* lacZ or trpE gene linked to RRV protein may be used to prepare polyclonal and monoclonal antibodies against this protein. Thereafter, these antibodies may be used to purify proteins by immunoaffinity chromatography, in diagnostic assays to quantitate the levels of protein and to localize proteins in tissues and individual cells by immunofluorescence and microscopy.

Intact native protein may also be produced in *E. coli* in large amounts for functional studies. Standard prokaryotic cloning vectors may also be used, for example pBR322, pUC18 or pUC19 as described in Sambrook et al. (Molecular Cloning: A Laboratory Manual, 2nd ed., vol. 1-3, Cold Spring Harbor, New York. 1989). Nucleic acids of RRV may be cloned into such vectors which may then be transformed into bacteria such as *E. coli* which may then be cultured so as to express the protein of interest. Other prokaryotic expression systems include, for instance, the arabinose-induced pBAD expression system that allows tightly controlled regulation of expression, the IPTG-induced pRSET system that facilitates rapid purification of recombinant proteins and the IPTG-induced pSE402 system that has been constructed for optimal translation of eukaryotic genes. These three systems are available commercially from Invitrogen and, when used according to the manufacturer's instructions, allow routine expression and purification of proteins.

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Methods and plasmid vectors for producing fusion proteins and intact native proteins in bacteria are described in Sambrook et al. (Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989, Chapter 17). Such fusion proteins may be made in large amounts, are easy to purify, and can be used to elicit antibody response. Native proteins can be produced in bacteria by placing a strong, regulated promoter and an efficient ribosome binding site upstream of the cloned gene. If low levels of protein are produced, additional steps may be taken to increase protein production; if high levels of protein are produced, purification is relatively easy. Suitable methods are presented in Sambrook et al. (Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989) and are well known in the art. Often, proteins expressed at high levels are found in insoluble inclusion bodies. Methods for extracting proteins from these aggregates are described by Sambrook et al. (Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989, Chapter 17).

Vector systems suitable for the expression of lacZ fusion genes include the pUR series of vectors (Ruther and Muller-Hill, 1983, *EMBO J.* 2:1791), pEX1-3 (Stanley and Luzio, 1984, *EMBO J.* 3:1429) and pMR100 (Gray et al., 1982, *Proc. Natl. Acad. Sci. USA* 79:6598). Vectors suitable for the production of intact native proteins include pKC30 (Shimatake and Rosenberg, 1981, *Nature* 292:128), pKK177-3 (Amann and Brosius, 1985, *Gene* 40:183) and pET-3 (Studiar and Moffatt, 1986, *J. Mol. Biol.* 189:113). The RRV fusion protein may be isolated from protein gels, lyophilized, ground into a powder and used as an antigen. The DNA sequence can also be transferred to other cloning vehicles, such as other plasmids, bacteriophages, cosmids, animal

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viruses and yeast artificial chromosomes (YACs) (Burke et al., 1987, Science 236:806-12). These vectors may then be introduced into a variety of hosts including somatic cells, and simple or complex organisms, such as bacteria, fungi (Timberlake and Marshall, 1989, Science 244:1313-7), invertebrates, plants (Gasser and Fraley, 1989, Science 244:1293), and mammals (Pursel et al., 1989, Science 244:1281-8), which cell or organisms are rendered transgenic by the introduction of one or more heterologous RRV DNAs.

Various yeast strains and yeast-derived vectors are commonly used for expressing and purifying proteins, for example, *Pichia pastoris* expression systems are available from Invitrogen (Carlsbad, CA). Such systems include suitable *Pichia pastoris* strains, vectors, reagents, transformants, sequencing primers and media.

Non-yeast eukaryotic vectors can also be used for expression of the RRV proteins. Examples of such systems are the well known Baculovirus system, the Ecdysone-inducible mammalian expression system that uses regulatory elements from *Drosophila melanogaster* to allow control of gene expression, and the Sindbis viral expression system that allows high level expression in a variety of mammalian cell lines. These expression systems are available from Invitrogen.

For expression in mammalian cells, the cDNA sequence may be ligated to heterologous promoters, such as the simian virus SV40, promoter in the pSV2 vector (Mulligan and Berg, 1981, *Proc. Natl. Acad. Sci. USA* 78:2072-6), and introduced into cells, such as monkey COS-1 cells (Gluzman, 1981, *Cell* 23:175-82), to achieve transient or long-term expression. The stable integration of the chimeric gene construct may be maintained in mammalian cells by biochemical selection, such as neomycin (Southern and Berg, 1982, *J. Mol. Appl. Genet.* 1:327-41) and mycophoenolic acid (Mulligan and Berg, 1981, *Proc. Natl. Acad. Sci. USA* 78:2072-6).

DNA sequences can be manipulated with standard procedures such as restriction enzyme digestion, fill-in with DNA polymerase, deletion by exonuclease, extension by terminal deoxynucleotide transferase, ligation of synthetic or cloned DNA sequences, site-directed sequence-alteration via single-stranded bacteriophage intermediate or with the use of specific oligonucleotides in combination with PCR.

The cDNA sequence (or portions derived from it) or a mini gene (a cDNA with an intron and its own promoter) may be introduced into eukaryotic expression vectors by conventional techniques. These vectors are designed to permit the transcription of the cDNA eukaryotic cells by providing regulatory sequences that initiate and enhance the transcription of the cDNA and ensure its proper splicing and polyadenylation. Vectors containing the promoter and enhancer regions of the SV40 or long terminal repeat (LTR) of the Rous Sarcoma virus and polyadenylation and splicing signal from SV40 are readily available (Mulligan and Berg, 1981, *Proc. Natl. Acad. Sci. USA* 78:2072-6; Gorman et al., 1982, *Proc. Natl. Acad. Sci USA* 78:6777-81). The level of expression of the cDNA can be manipulated with this type of vector, either by using promoters

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that have different activities (for example, the baculovirus pAC373 can express cDNAs at high levels in *S. frugiperda* cells (Summers and Smith, 1985, Genetically Altered Viruses and the Environment, Fields et al. (Eds.) 22:319-328, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York.) or by using vectors that contain promoters amenable to modulation, for example, the glucocorticoid-responsive promoter from the mouse mammary tumor virus (Lee et al., 1982, Nature 294:228). The expression of the cDNA can be monitored in the recipient cells 24 to 72 hours after introduction (transient expression).

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In addition, some vectors contain selectable markers such as the *gpt* (Mulligan and Berg, 1981, *Proc. Natl. Acad. Sci. USA* 78:2072-6) or *neo* (Southern and Berg, 1982, *J. Mol. Appl. Genet.* 1:327-41) bacterial genes. These selectable markers permit selection of transfected cells that exhibit stable, long-term expression of the vectors (and therefore the cDNA). The vectors can be maintained in the cells as episomal, freely replicating entities by using regulatory elements of viruses such as papilloma (Sarver et al., 1981, *Mol. Cell Biol.* 1:486) or Epstein-Barr (Sugden et al., 1985, *Mol. Cell Biol.* 5:410). Alternatively, one can also produce cell lines that have integrated the vector into genomic DNA. Both of these types of cell lines produce the gene product on a continuous basis. One can also produce cell lines that have amplified the number of copies of the vector (and therefore of the cDNA as well) to create cell lines that can produce high levels of the gene product (Alt et al., 1978, *J. Biol. Chem.* 253:1357).

The transfer of DNA into eukaryotic, in particular human or other mammalian cells, is now a conventional technique. The vectors are introduced into the recipient cells as pure DNA (transfection) by, for example, precipitation with calcium phosphate (Graham and vander Eb, 1973, Virology 52:466) or strontium phosphate (Brash et al., 1987, Mol. Cell Biol. 7:2013), electroporation (Neumann et al., 1982, EMBO J. 1:841), lipofection (Felgner et al., 1987, Proc. Natl. Acad. Sci USA 84:7413), DEAE dextran (McCuthan et al., 1968, J. Natl. Cancer Inst. 41:351), microinjection (Mueller et al., 1978, Cell 15:579), protoplast fusion (Schafner, 1980, Proc. Natl. Acad. Sci. USA 77:2163-7), or pellet guns (Klein et al., 1987, Nature 327:70). Alternatively, the cDNA can be introduced by infection with virus vectors. Systems are developed that use, for example, retroviruses (Bernstein et al., 1985, Gen. Engrg. 7:235), adenoviruses (Ahmad et al., 1986, J. Virol. 57:267), or Herpes virus (Spaete et al., 1982, Cell 30:295).

These eukaryotic expression systems can be used for studies of RRV genes and mutant forms of these genes, the RRV proteins and mutant forms of these proteins. Such uses include, for example, the identification of regulatory elements located in the 5' region of RRV genes on genomic clones that can be isolated from genomic DNA libraries, such as human or mouse libraries, using the information contained in the present invention. The eukaryotic expression systems may also be used to study the function of the normal complete protein, specific portions of the protein, or of naturally occurring or artificially produced mutant proteins. Naturally occurring RRV wild-type or mutant proteins may exist in a variety of cancers or diseases, while artificially

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produced mutant proteins can be designed by site directed mutagenesis as described above. These latter studies may probe the function of any desired amino acid residue in the protein by mutating the nucleatide coding for that amino acid.

Using the above techniques, the expression vectors containing RRV genes or cDNA sequence or fragments or variants or mutants thereof can be introduced into human cells, mammalian cells from other species or non-mammalian cells as desired. The choice of cell is determined by the purpose of the treatment. For example, monkey COS cells (Gluzman, 1981, Cell 23:175-82) that produce high levels of the SV40 T antigen and permit the replication of vectors containing the SV40 origin of replication may be used. Similarly, Chinese hamster ovary (CHO), mouse NIH 3T3 fibroblasts or human fibroblasts or lymphoblasts may be used.

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One method that can be used to express RRV polypeptides from the cloned RRV cDNA sequence in mammalian cells is to use the cloning vector, pXTI. This vector is commercially available from Stratagene, contains the Long Terminal Repeats (LTRs) and a portion of the GAG gene from Moloney Murine Leukemia Virus. The position of the viral LTRs allows highly efficient, stable transfection of the region within the LTRs. The vector also contains the Herpes Simplex Thymidine Kinase promoter (TK), active in embryonal cells and in a wide variety of tissues in mice, and a selectable neomycin gene conferring G418 resistance. Two unique restriction sites BgIII and XhoI are directly downstream from the TK promoter. RRV cDNA, including the entire open reading frame for an RRV protein such as IL-6 and the 3' untranslated region of the cDNA is cloned into one of the two unique restriction sites downstream from the promoter.

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc.) under conditions outlined in the product specification. Positive transfectants are selected after growing the transfected cells in 600 µg/ml G418 (Sigma, St. Louis, MO). The protein is released into the supernatant and may be purified by standard immunoaffinity chromatography techniques using antibodies raised against RRV proteins (see Example18).

Expression of RRV proteins in eukaryotic cells can be used as a source of proteins to raise antibodies. The RRV proteins may be extracted following release of the protein into the supernatant as described above, or, the cDNA sequence may be incorporated into a eukaryotic expression vector and expressed as a chimeric protein with, for example, β -globin. Antibody to β -globin is thereafter used to purify the chimeric protein. Corresponding protease cleavage sites engineered between the β -globin gene and the cDNA are then used to separate the two polypeptide fragments from one another after translation. One useful expression vector for generating β -globin chimeric proteins is pSG5 (Stratagene). This vector encodes rabbit β -globin.

The present invention thus encompasses recombinant vectors which comprise all or part of RRV genome or cDNA sequences, for expression in a suitable host. The RRV DNA is operatively linked in the vector to an expression control sequence in the recombinant DNA molecule so that a

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RRV polypeptide can be expressed. The expression control sequence may be selected from the group consisting of sequences that control the expression of genes of prokaryotic or eukaryotic cells and their viruses and combinations thereof. The expression control sequence may be specifically selected from the group consisting of the lac system, the trp system, the tac system, the trc system, major operator and promoter regions of phage lambda, the control region of fd coat protein, the early and late promoters of SV40, promoters derived from polyoma, adenovirus, retrovirus, baculovirus and simian virus, the promoter for 3-phosphoglycerate kinase, the promoters of yeast acid phosphatase, the promoter of the yeast alpha-mating factors and combinations thereof.

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The host cell, which may be transfected with the vector of this invention, may be selected from the group consisting of: *E. coli*, *Pseudomonas*, *Bacillus subtilis*, *Bacillus stearothermophilus* or other bacilli; other bacteria; yeast; fungi; plant; insect; mouse or other animal; or human tissue cells.

It is appreciated that for mutant or variant RRV DNA sequences, similar systems are employed to express and produce the mutant or variant product.

EXAMPLE 18

Production of Antibodies to RRV and RRV Proteins

Polyclonal or monoclonal antibodies (including humanized monoclonal antibodies) and fragments of monoclonal antibodies such as Fab, F(ab')2 and Fv fragments, as well as any other agent capable of specifically binding to an RRV protein, may be produced to the RRV virion or any of the RRV proteins (for example odd-numbered SEQ ID Nos 3-165). Optimally, antibodies raised against an RRV protein would specifically detect the RRV protein of interest (or a virion containing the protein of interest). That is, such antibodies would recognize and bind the protein and would not substantially recognize or bind to other proteins found in human or other cells. The determination that an antibody specifically detects the RRV protein is made by any one of a number of standard immunoassay methods; for instance, the Western blotting technique (Sambrook et al., 1989, Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Laboratory, Cold Spring Harbor, New York).

To determine that a given antibody preparation (such as one produced in a mouse) specifically detects the RRV protein by Western blotting, total cellular protein is extracted from murine myeloma cells and electrophoresed on a SDS-polyacrylamide gel. The proteins are then transferred to a membrane (for example, nitrocellulose) by Western blotting, and the antibody preparation is incubated with the membrane. After washing the membrane to remove non-specifically bound antibodies, the presence of specifically bound antibodies is detected by the use of an anti-mouse antibody conjugated to an enzyme such as alkaline phosphatase; application of the substrate 5-bromo-4-chloro-3-indolyl phosphate/nitro blue tetrazolium results in the production of

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a dense blue compound by immuno-localized alkaline phosphatase. Antibodies which specifically detect an RRV protein will, by this technique, be shown to bind to the RRV protein band (which will be localized at a given position on the gel determined by its molecular weight). Non-specific binding of the antibody to other proteins (such as serum albumin) may occur and may be detectable as a weak signal on the Western blot. The non-specific nature of this binding will be recognized by one skilled in the art by the weak signal obtained on the Western blot relative to the strong primary signal arising from the specific antibody-VIAP protein binding.

A substantially pure virion can be obtained, or substantially pure RRV protein suitable for use as an immunogen is isolated by purification or recombinant expression. Concentration of protein in the final preparation is adjusted, for example, by concentration on an Amicon filter device, to the level of a few micrograms per milliliter. Monoclonal or polyclonal antibody to the protein can then be prepared as described by Harlow and Lane (Antibodies, A Laboratory Manual, Cold Spring Harbor Press. 1988).

Alternatively, antibodies may be raised against synthetic RRV peptides synthesized on a commercially available peptide synthesizer (see Example 26) based upon the predicted amino acid sequence of the RRV protein (Harlow and Lane, Antibodies, A Laboratory Manual, Cold Spring Harbor Press. 1988).

Another method of raising antibodies against RRV proteins is by subcutaneous injection of a DNA vector which expresses the RRV protein into laboratory animals, such as mice. Delivery of the recombinant vector into the animals may be achieved using a hand-held form of the Biolistic system (Sanford et al., 1987, *Particulate Sci. Technol.* 5:27-37) as described by Tang et al. (*Nature* 356:152-4, 1992). Expression vectors suitable for this purpose may include those which express the RRV protein under the transcriptional control of either the human β -actin promoter or the cytomegalovirus (CMV) promoter.

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Monoclonal Antibody Production by Hybridoma Fusion

Monoclonal antibody to epitopes of the RRV protein identified and isolated as described can be prepared from murine hybridomas according to the classical method of Kohler and Milstein (Nature 256:495, 1975) or derivative methods thereof. Briefly, a mouse is repetitively inoculated with a few micrograms of the selected protein over a period of a few weeks. The mouse is then sacrificed, and the antibody-producing cells of the spleen isolated. The spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as ELISA, as originally described by Engvall (Enzymol. 70:419, 1980), and derivative methods thereof.

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Selected positive clones can be expanded and their monoclonal antibody product harvested for use.

Detailed procedures for monoclonal antibody production are described in Harlow and Lane

(Antibodies: A Laboratory Manual. 1988, Cold Spring Harbor Laboratory, New York).

Polyclonal Antibody Production by Immunization

Polyclonal antiserum containing antibodies to heterogeneous epitopes of a single protein can be prepared by immunizing suitable animals with the expressed protein (for example see Example 17), which can be unmodified or modified to enhance immunogenicity. Effective polyclonal antibody production is affected by many factors related both to the antigen and the host species. For example, small molecules tend to be less immunogenic than others and may require the use of carriers and adjuvant. Also, host animals vary in response to site of inoculations and dose, with both inadequate or excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. An effective immunization protocol for rabbits can be found in Vaitukaitis et al. (*J. Clin. Endocrinol. Metab.* 33:988-91, 1971).

Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against known concentrations of the antigen, begins to fall. See, for example, Ouchterlony et al. (In: Handbook of Experimental Immunology, Wier, D. (ed.). Chapter 19. Blackwell. 1973). Plateau concentration of antibody is usually in the range of 0.1 to 0.2 mg/ml of serum (about 12 μ M). Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as described, for example, by Fisher (Manual of Clinical Immunology, Chapter 42. 1980).

25 Labeled Antibodies

Antibodies of the present invention can be conjugated with various labels for their direct detection (see Chapter 9, Harlow and Lane, Antibodies: A Laboratory Manual. 1988). The label, which may include, but is not limited to, a radiolabel, enzyme, fluorescent probe, or biotin, is chosen based on the method of detection available to the user.

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EXAMPLE 19

Diagnostic Methods

An embodiment of the present invention is a method for screening a subject to determine if the subject has been infected with RRV. One major application of the RRV sequence information presented herein is in the area of diagnostic testing for predisposition to a disease (such as Kaposi's Sarcoma and lymphoproliferative disorders) that develops in at least a sub-set of hosts infected with RRV. The gene sequence of the RRV genes, including intron-exon boundaries

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is also useful in such diagnostic methods. The method includes providing a biological sample obtained from the subject, in which sample includes DNA or RNA, and providing an assay for detecting in the biological sample the presence of any of the RRV genes or proteins. Suitable biological samples include samples obtained from body cells, such as those present in peripheral blood, urine, saliva, tissue biopsy, surgical specimen, fine needle aspirate specimen, amniocentesis samples and autopsy material. The detection in the biological sample may be performed by a number of methodologies, as outlined below.

The foregoing assay may be assembled in the form of a diagnostic kit and preferably comprises either: hybridization with oligonucleotides; PCR amplification of the gene or a part thereof using oligonucleotide primers; RT-PCR amplification of the RNA or a part thereof using oligonucleotide primers; or direct sequencing of any of the RRV genes present in a subject using oligonucleotide primers. The efficiency of these molecular genetic methods should permit the rapid identification of patients infected with RRV.

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One embodiment of such detection techniques is the polymerase chain reaction amplification of reverse transcribed RNA (RT-PCR) of RNA isolated from cells (for example lymphocytes) followed by direct DNA sequence determination of the products. The presence of one or more RRV genes is taken as indicative of a potential RRV infection.

Alternatively, DNA extracted from lymphocytes or other cells may be used directly for amplification. The direct amplification from genomic DNA would be appropriate for analysis of an entire RRV gene including regulatory sequences located upstream and downstream from the open reading frame. Recent reviews of direct DNA diagnosis have been presented by Caskey (Science 236:1223-1228, 1989) and by Landegren et al. (Science 242:229-37, 1989).

Further studies of RRV genes isolated from subjects may reveal particular mutations, deletions or alterations in gene sequences, which occur at a high frequency within particular populations of individuals. In this case, rather than sequencing the entire RRV gene, it may be possible to design DNA diagnostic methods to specifically detect the most common RRV mutations, deletions or alterations in gene sequences.

The detection of specific DNA mutations or alterations in gene sequences may be achieved by methods such as hybridization using specific oligonucleotides (Wallace et al., 1986, Cold Spring Harbor Symp. Quant. Biol. 51:257-61), direct DNA sequencing (Church and Gilbert, 1984, Proc. Natl. Acad. Sci. USA. 81:1991-5), the use of restriction enzymes (Flavell et al., 1978, Cell 15:25; Geever et al., 1981, Proc. Natl. Acad. Sci USA 78:5081), discrimination on the basis of electrophoretic mobility in gels with denaturing reagent (Myers and Maniatis, 1986, Cold Spring Harbor Symp. Quant. Biol. 51:275-284), RNase protection (Myers et al., 1985, Science 230:1242), chemical cleavage (Cotton et al., 1985, Proc. Natl. Acad. Sci. USA 85:4397-401), and the ligase-mediated detection procedure (Landegren et al., 1988, Science 241:1077).

Oligonucleotides specific to normal, mutant or alterative sequences are chemically

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synthesized using commercially available machines, labeled radioactively with isotopes (such as ³²P) or non-radioactively, with tags such as biotin (Ward and Langer et al., 1981. *Proc. Natl. Acad. Sci. USA* 78:6633-57), and hybridized to individual DNA samples immobilized on membranes or other solid supports by dot-blot or transfer from gels after electrophoresis. The presence of these specific sequences are visualized by methods such as autoradiography or fluorometric (Landegren et al., 1989, *Science* 242:229-37) or colorimetric reactions (Gebeyehu et al., 1987, *Nucleic Acids Res.* 15:4513-34). The absence of hybridization would indicate a mutation in the particular region of the gene, or that the patient is not infected with RRV.

Sequence differences between disclosed and other forms of RRV genes may also be revealed by the direct DNA sequencing method of Church and Gilbert (*Proc. Natl. Acad. Sci. USA* 81:1991-5, 1988). Cloned DNA segments may be used as probes to detect specific DNA segments. The sensitivity of this method is greatly enhanced when combined with PCR (Wrichnik et al., 1987, *Nucleic Acids Res.* 15:529-42; Wong et al., 1987, *Nature* 330:384-6; Stoflet et al., 1988, *Science* 239:491-4). In this approach, a sequencing primer which lies within the amplified sequence is used with double-stranded PCR product or single-stranded template generated by a modified PCR. The sequence determination is performed by conventional procedures with radiolabeled nucleotides or by automatic sequencing procedures with fluorescent tags.

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Sequence alterations may occasionally generate fortuitous restriction enzyme recognition sites or may eliminate existing restriction sites. Changes in restriction sites are revealed by the use of appropriate enzyme digestion followed by conventional gel-blot hybridization (Southern, 1975, *J. Mol. Biol.* 98:503). DNA fragments carrying the site (either normal, mutant, or alternative) are detected by their reduction in size or increase of corresponding restriction fragment numbers. Genomic DNA samples may also be amplified by PCR prior to treatment with the appropriate restriction enzyme; fragments of different sizes are then visualized under UV light in the presence of ethidium bromide after gel electrophoresis.

Screening based on DNA sequence differences may be achieved by detection of alteration in electrophoretic mobility of DNA fragments in gels with or without denaturing reagent. Small sequence deletions and insertions can be visualized by high-resolution gel electrophoresis. For example, a PCR product with small deletions is clearly distinguishable from a normal sequence on an 8% non-denaturing polyacrylamide gel (WO 91/10734; Nagamine et al., 1989, Am. J. Hum. Genet. 45:337-9). DNA fragments of different sequence compositions may be distinguished on denaturing formamide gradient gels in which the mobilities of different DNA fragments are retarded in the gel at different positions according to their specific "partial-melting" temperatures (Myers et al., 1985, Science 230:1242). Alternatively, a method of detecting a mutation comprising a single base substitution or other small change could be based on differential primer length in a PCR. For example, an invariant primer could be used in addition to a primer specific for a mutation. The PCR products of the normal and mutant genes can then be differentially

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detected in acrylamide gels.

In addition to conventional gel-electrophoresis and blot-hybridization methods, DNA fragments may also be visualized by methods where the individual DNA samples are not immobilized on membranes. The probe and target sequences may be both in solution, or the probe sequence may be immobilized (Saiki et al., 1989, *Proc. Nat. Acad. Sci. USA* 86:6230-4). A variety of detection methods, such as autoradiography involving radioisotopes, direct detection of radioactive decay (in the presence or absence of scintillant), spectrophotometry involving calorigenic reactions and fluorometry involved fluorogenic reactions, may be used to identify specific individual genotypes.

If more than one mutation or alternative sequence is frequently encountered in one or more RRV genes, a system capable of detecting such multiple mutations would be desirable. For example, a PCR with multiple, specific oligonucleotide primers and hybridization probes may be used to identify all possible mutations or alternative sequences at the same time (Chamberlain et al., 1988, *Nucl. Acids Res.* 16:1141-55). The procedure may involve immobilized sequence-specific oligonucleotides probes (Saiki et al., 1989, *Proc. Nat. Acad. Sci. USA* 86:6230-4).

EXAMPLE 20

Quantitation of RRV Proteins

An alternative method of determining if a subject has been infected with RRV is to quantitate the level of one or more RRV proteins in the cells of a subject. This diagnostic tool would be useful for detecting the levels of RRV proteins which result from, for example, infection by RRV. These diagnostic methods, in addition to those described in EXAMPLE 19, provide an enhanced ability to diagnose susceptibility to diseases caused by RRV infection.

The determination of RRV protein levels would be an alternative or supplemental approach to the direct determination of the presence of one or more RRV genes by the methods outlined above in EXAMPLE 19. The availability of antibodies specific to one or more of the RRV proteins (for example those described in Example 18) will facilitate the quantitation of cellular RRV proteins by one of a number of immunoassay methods which are well known in the art and are presented in Harlow and Lane (Antibodies, A Laboratory Manual, Cold Spring Harbor Laboratory, New York. 1988).

Such assays permit both the detection of RRV proteins in a biological sample and the quantitation of such proteins. Typical methods involve: providing a biological sample of the subject in which the sample contains cellular proteins, and providing an immunoassay for quantitating the level of RRV protein in the biological sample. This can be achieved by combining the biological sample with an RRV specific binding agent, such as an anti-RRV antibody (such as monoclonal or polyclonal antibodies), so that complexes form between the binding agent and the RRV protein present in the sample, and then detecting or quantitating such complexes.

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In particular forms, these assays may be performed with the RRV specific binding agent immobilized on a support surface, such as in the wells of a microtiter plate or on a column. The biological sample is then introduced onto the support surface and allowed to interact with the specific binding agent so as to form complexes. Excess biological sample is then removed by washing, and the complexes are detected with a reagent, such as a second anti- RRV protein antibody that is conjugated with a detectable marker.

In an alternative assay, the cellular proteins are isolated and subjected to SDS-PAGE followed by Western blotting, for example as described in Example 18. After resolving the proteins, the proteins are transferred to a membrane, which is probed with specific binding agents that recognize any of the RRV proteins. The proteins are detected, for example with HRP-conjugated secondary antibodies, and quantitated.

In yet another assay, the level of one or more RRV proteins in cells is analyzed using microscopy. Using specific binding agents which recognize RRV, samples can be analyzed for the presence of one or more RRV proteins. For example, frozen biopsied tissue sections are thawed at room temperature and fixed with acetone at -200°C for 5 minutes. Slides are washed twice in cold PBS for 5 minutes each, then air-dried. Sections are covered with 20-30 μ l of antibody solution (15-45 μ g/ml) (diluted in PBS, 2% BSA at 15-50 μ g/ml) and incubated at room temperature in humidified chamber for 30 minutes. Slides are washed three times with cold PBS 5 minutes each, allowed to air-dry briefly (5 minutes) before applying 20-30 μ l of the second antibody solution (diluted in PBS, 2% BSA at 15-50 μ g/ml) and incubated at room temperature in humidified chamber for 30 minutes. The label on the second antibody may contain a fluorescent probe, enzyme, radiolabel, biotin, or other detectable marker. The slides are washed three times with cold PBS 5 minutes each then quickly dipped in distilled water, air-dried, and mounted with PBS containing 30% glycerol. Slides can be stored at 4°C prior to viewing.

For samples prepared for electron microscopy (versus light microscopy), the second antibody is conjugated to gold particles. Tissue is fixed and embedded with epoxy plastics, then cut into very thin sections ($\sim 1-2~\mu m$). The specimen is then applied to a metal grid, which is then incubated in the primary anti-RRV antibody, washed in a buffer containing BSA, then incubated in a secondary antibody conjugated to gold particles (usually 5-20 nm). These gold particles are visualized using electron microscopy methods.

For the purposes of quantitating the RRV proteins, a biological sample of the subject, which sample includes cellular proteins, is required. Such a biological sample may be obtained from body cells, such as those present in which expression of the protein has been detected. The expression of RRV proteins in peripheral blood leukocytes is clearly the most accessible and convenient source from which specimens can be obtained. Specimens can be obtained from peripheral blood, urine, saliva, tissue biopsy, amniocentesis samples, surgical specimens, fine needle aspirates, and autopsy material, particularly cancer cells. Quantitation of RRV proteins

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would be made by immunoassay and compared to levels of the protein found in non-RRV expressing cells or to the level of RRV proteins in non-RRV infected cells (cells of the same origin that are not infected). A significant (preferably 50% or greater) increase in the amount of one or more RRV proteins in the cells of a subject compared to the amount of one or more RRV proteins found in non-RRV infected cells or that found in normal cells, would be taken as an indication that the subject may have been infected with RRV.

EXAMPLE 21

Sequence Variants of RRV

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The amino acid sequence of the RRV proteins which are encoded by the RRV cDNAs (even-numbered SEQ ID NOS 2-164), are shown in odd-numbered SEQ ID NOS 3-165. Having presented the nucleotide sequence of the RRV genome and cDNAs and the amino acid sequence of these proteins, this invention now also facilitates the creation of DNA molecules, and thereby proteins, which are derived from those disclosed but which vary in their precise nucleotide or amino acid sequence from those disclosed. Such variants may be obtained through a combination of standard molecular biology laboratory techniques and the nucleotide sequence information disclosed by this invention.

Variant DNA molecules include those created by standard DNA mutagenesis techniques, for example, M13 primer mutagenesis. Details of these techniques are provided in Sambrook et al. (In: Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989, Chapter 15). By the use of such techniques, variants may be created which differ in minor ways from those disclosed. DNA molecules and nucleotide sequences which are derivatives of those specifically disclosed herein and which differ from those disclosed by the deletion, addition or substitution of nucleotides while still encoding a protein which possesses the functional characteristics of the RRV proteins are comprehended by this invention. Also within the scope of this invention are small DNA molecules which are derived from the disclosed DNA molecules. Such small DNA molecules include oligonucleotides suitable for use as hybridization probes or polymerase chain reaction (PCR) primers. As such, these small DNA molecules will comprise at least a segment of the RRV cDNA molecules or the RRV gene and, for the purposes of PCR, will comprise at least a 15 or a 20-50 nucleotide sequence of the RRV cDNAs (even-numbered SEQ ID Nos 2-164) or the RRV genes (i.e., at least 20-50 consecutive nucleotides of the RRV cDNA or gene sequences). DNA molecules and nucleotide sequences which are derived from the disclosed DNA molecules as described above may also be defined as DNA sequences which hybridize under

stringent conditions to the DNA sequences disclosed, or fragments thereof.

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Hybridization conditions resulting in particular degrees of stringency will vary depending upon the nature of the hybridization method of choice and the composition and length of the hybridizing DNA used. Generally, the temperature of hybridization and the ionic strength (especially the Na+ concentration) of the hybridization buffer will determine the stringency of hybridization. Calculations regarding hybridization conditions required for attaining particular degrees of stringency are discussed by Sambrook et al. (In: Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989 ch. 9 and 11), herein incorporated by reference. By way of illustration only, a hybridization experiment may be performed by hybridization of a DNA molecule (for example, a deviation of the RRV cDNA) to a target DNA molecule (for example, the RRV cDNA) which has been electrophoresed in an agarose gel and transferred to a nitrocellulose membrane by Southern blotting (Southern, J. Mol. Biol. 98:503, 1975), a technique well known in the art and described in Sambrook et al. (Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989). Hybridization with a target probe labeled with [32P]-dCTP is generally carried out in a solution of high ionic strength such as 6xSSC at a temperature that is 20-25°C below the melting temperature, Tm, described below. For such Southern hybridization experiments where the target DNA molecule on the Southern blot contains 10 ng of DNA or more, hybridization is typically carried out for 6-8 hours using 1-2 ng/ml radiolabeled probe (of specific activity equal to 109 CPM/µg or greater). Following hybridization, the nitrocellulose filter is washed to remove background hybridization. The washing conditions should be as stringent as possible to remove background hybridization but to retain a specific hybridization signal. The term Tm represents the temperature above which, under the prevailing ionic conditions, the radiolabeled probe molecule will not hybridize to its target DNA molecule. The Tm of such a hybrid molecule may be estimated from the following equation (Bolton and McCarthy, Proc. Natl. Acad. Sci. USA 48:1390, 1962): Tm = 81.5°C - 16.6(log10[Na+]) + 0.41(%G+C) - 0.63(% formamide) - (600/1); where l = the length of the hybrid in base pairs.

This equation is valid for concentrations of Na⁺ in the range of 0.01 M to 0.4 M, and it is less accurate for calculations of T_m in solutions of higher [Na⁺]. The equation is also primarily valid for DNAs whose G+C content is in the range of 30% to 75%, and it applies to hybrids greater than 100 nucleotides in length (the behavior of oligonucleotide probes is described in detail in Ch. 11 of Sambrook et al. (Molecular Cloning: A Laboratory Manual, Cold Spring Harbor, New York, 1989).

Thus, by way of example, for a 150 base pair DNA probe derived from the open reading frame of the RRV cDNA (with a hypothetical %GC = 45%), a calculation of hybridization conditions required to give particular stringencies may be made as follows: For this example, it is assumed that the filter will be washed in 0.3 xSSC solution following hybridization, thereby: $[Na^+] = 0.045M$; %GC = 45%; Formamide concentration = 0; I = 150 base pairs; $I_m = 81.5$ –

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 $16.6(\log_{10}[Na^+]) + (0.41 \times 45) - (600/150)$; and so $T_m = 74.4$ °C.

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The T_m of double-stranded DNA decreases by 1-1.5°C with every 1% decrease in homology (Bonner et al., *J. Mol. Biol.* 81:123, 1973). Therefore, for this given example, washing the filter in 0.3 xSSC at 59.4-64.4°C will produce a stringency of hybridization equivalent to 90%; that is, DNA molecules with more than 10% sequence variation relative to the target RRV cDNA will not hybridize. Alternatively, washing the hybridized filter in 0.3 xSSC at a temperature of 65.4-68.4°C will yield a hybridization stringency of 94%; that is, DNA molecules with more than 6% sequence variation relative to the target RRV cDNA molecule will not hybridize. The above example is given entirely by way of theoretical illustration. One skilled in the art will appreciate that other hybridization techniques may be utilized and that variations in experimental conditions will necessitate alternative calculations for stringency.

In particular embodiments of the present invention, stringent conditions may be defined as those under which DNA molecules with more than 25%, 15%, 10%, 6% or 2% sequence variation (also termed "mismatch") will not hybridize.

The degeneracy of the genetic code further widens the scope of the present invention as it enables major variations in the nucleotide sequence of a DNA molecule while maintaining the amino acid sequence of the encoded protein. For example, the eleventh amino acid residue of the RRV MIP protein is alanine (SEQ ID NO 25). This is encoded in the RRV cDNA by the nucleotide codon triplet GCG. Because of the degeneracy of the genetic code, three other nucleotide codon triplets, GCT, GCA and GCC, also code for alanine. Thus, the nucleotide sequence of the RRV DNA could be changed at this position to any of these three codons without affecting the amino acid composition of the encoded protein or the characteristics of the protein. Based upon the degeneracy of the genetic code, variant DNA molecules may be derived from the DNA molecules disclosed herein using standard DNA mutagenesis techniques as described above, or by synthesis of DNA sequences. DNA sequences which do not hybridize under stringent conditions to the DNA sequences disclosed by virtue of sequence variation based on the degeneracy of the genetic code are herein also comprehended by this invention.

The invention also includes DNA sequences that are substantially identical to any of the DNA sequences disclosed herein, where substantially identical means a sequence that has identical nucleotides in at least 75%, 80%, 85%, 90%, 95%, 98%, or even 99% of the aligned sequences.

One skilled in the art will recognize that the DNA mutagenesis techniques described above may be used not only to produce variant DNA molecules, but will also facilitate the production of proteins which differ in certain structural aspects from the RRV proteins, yet which proteins are clearly derivative of this protein and which maintain the essential characteristics of the RRV proteins. Newly derived proteins may also be selected in order to obtain variations on the characteristic of the RRV proteins, as described above. Such derivatives include those with variations in amino acid sequence including minor deletions, additions and substitutions.

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While the site for introducing an amino acid sequence variation is predetermined, the mutation per se need not be predetermined. For example, in order to optimize the performance of a mutation at a given site, random mutagenesis may be conducted at the target codon or region and the expressed protein variants screened for the optimal combination of desired activity. Techniques for making substitution mutations at predetermined sites in DNA having a known sequence as described above are well known.

Amino acid substitutions are typically of single residues; insertions usually will be on the order of about from 1 to 10 amino acid residues; and deletions will range about from 1 to 30 residues. Deletions or insertions preferably are made in adjacent pairs, i.e., a deletion of two residues or insertion of two residues. Substitutions, deletions, insertions or any combination thereof may be combined to arrive at a final construct. Obviously, the mutations that are made in the DNA encoding the protein must not place the sequence out of reading frame and preferably will not create complementary regions that could produce secondary mRNA structure.

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Substitutional variants are those in which at least one residue in the amino acid sequence has been removed and a different residue inserted in its place. Such substitutions generally are made conservatively, as defined above.

The effects of these amino acid substitutions or deletions or additions may be assessed for derivatives of the RRV proteins by assays in which DNA molecules encoding the derivative proteins are transfected into cells using routine procedures. These RRV proteins are expressed recombinantly (for example see Example 17), purified, and analyzed for their ability to cause symptoms associated with RRV infection, for example KS-like symptoms in rhesus macaque monkeys, as described in Examples 13 and 23.

EXAMPLE 22

Cloning RRV in Other Species

Having presented the genomic (SEQ ID NO 1) and cDNA nucleotide sequences of the rhesus macaque RRV (even-numbered SEQ ID Nos 2-164) and the amino acid sequence of the encoded proteins (odd-numbered SEQ ID Nos 3-165), this invention now also facilitates the identification of DNA molecules, and thereby proteins, which are the RRV homologs in other species. These other homologs can be derived from those sequences disclosed, but which vary in their precise nucleotide or amino acid sequence from those disclosed. Such variants may be obtained through a combination of standard molecular biology laboratory techniques and the nucleotide and amino acid sequence information disclosed by this invention.

The Japanese macaque RRV isolate was isolated from a lesion that was minced and cocultured with primary rhesus fibroblasts. The isolate was then cloned by limiting dilution and a stock of virus generated from this clone. Total cellular DNA was harvested from virus infected cells and the DNA subjected to degenerate PCR for viral DNA polymerase, exactly as described - 56 -

above for RRV. Once confirmed, a cosmid library of this virus was made from purified viral DNA (as described for RRV) and then a portion of the protein genes was cloned and sequenced.

Results for this analysis are shown in the following Table 1:

TABLE 1

5 RRV Sequences from Japanese Macaque

Total number of amino acid residues inferred: 972 Number of differences compared to RRV: 29

Percent identity: 97.02%

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Japanese Macaque Data

These are end sequences. For ORFs represented twice, section I is from one plasmid, section II is from another plasmid. These are non-overlapping sections.

Orf 7 section I
GLFNSIDDTINALSRDCSVTFFQQANYTNVMRKQNELFTRLNSILCQGSAGSQKPATPSEPRT
ATVAATAASDVIKDAQYRKEQYMKKVARDGFKKLTECLQTQSAVLANALCMRVWGGVA
YGEASELVNHFLLRRFVALPWEARCRSNQILFENSKYIKNSLYSQRLSREHVEIITLQFYGLI
TGPLTRQSDLFPGPANVVLAQCFEAAGMLPHHKMLVSEMIW

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Orf 7 section II

PIESLFCGGLFNSIDDTINALSRDCSVTFFQQANYTNXMRKQNELFTRLNSILCQGSAGSXKP ATPSEPRTATVXATAASDVIKDAQYRXEQYMKKVARDXFKKLTECLQTQSAVLANALCMR RMGGRRI

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Orf 8

YRKVATSVTVYRGWTETAVTGKQEVIRPVPQYEINHMDTTYQCFSSMRVNVNGIENTYTD RDFTNQTVFLQPVEGLTDNIQRYFSQPVLYTTPGWFPGIYRVRTTVNCEIVDMIARSAEPYS YFVTALGDTVEVSPFCLNDSTCSVADKAENGLGVRVLTNYTIVDFATRTPTTETRVFADSGE

30 YTVSWKAEDPKSAVCALTLWKTFPRAIOTTHESOLPLCGORR

Orf 9 section I

VPSRFQTDIIPSGTVLKLLGRTENGTSVCVNVFRQQVYFYAKVPAGVNVTHVLQQALKNTA GRAACGFSTRRVTKKILKTYDVAEHPVTEITLSSGSMLSTLSDRLVACGCEVFESNVDAVRR

35 FVLDHGFTTFGWYSCARATPRLAXRDARTALEFDCSWEDLSV

Orf 9 section II

MDFFNPYLGPRGPRPPSHKCTDAPAPAGAVQPPPDVCRLIPACLRTPGAGGMIPVTIPFPPTY FENGARGDVLLAHERSMWTARGQRPVVPDPQDQSITFHAYDVVETTYAADRCAEV

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Orf 10

AQMKIIYAPGDPNAEIVLGQSGPVLPTHTGGRVLGVYADAEKTIQPGSSAEVRVQLIFPTGSA ARGDLAFLVAGVAPEPLFIVTPTLLLSGCTTHLRLFNPNGT

45 Orf 29b

NVAVEGNSSQDAGVAIATVLNEICSVPLSFLHHADKNTLIRSPIYMLGPEKAKAFESFIYALN SGTFSASQTVVSHTIKLSFDPVAYLIDQIKAIRCIPLKDGGHTYCAKQKTMSDDVLVATVMA HYMATNDKFVFKSLE

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EXAMPLE 23

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The present disclosure provides a virus that is involved in the causation or progression of certain diseases, such as KS, and therefore provides an animal model and assays directed to identifying potential pharmaceutical agents that inhibit the biological activity of the virus. Drug screening assays which determine whether or not a drug has activity against the virus can include incubating a compound to be evaluated for use in treatment of the condition with cells which express the RRV proteins or peptides, and determining the effect of the compound on the activity of the virus. In vitro assays in which the virus is maintained in suitable cell culture are preferred, though in vivo animal models would also be effective.

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In vitro assays include infecting cells such as rhesus fibroblasts, peripheral blood leukocytes or susceptible T cell lines such as MT-4 with the agent of interest in the presence of varying concentrations of compounds targeted against viral replication, including nucleoside analogs, chain terminators, antisense oligonucleotides and random polypeptides. (Asada et al., *J. Clin. Microbiol.* 27:2204, 1989; Kikuta et al., Lancet 7:861, 1989). Infected cultures and their supernatants can be assayed for the total amount of virus, including the presence of the viral genome, by quantitative PCR, by dot blot assays, or by using immunologic methods. For example, a culture of susceptible cells could be infected with the RRV in the presence of various concentrations of drug, fixed on slides after a period of days, and examined for viral antigen by indirect immunofluorescence with monoclonal antibodies to viral polypeptides (Kikuta et al, supra). Alternatively, chemically adhered MT-4 cell monolayers can be used for an infectious agent assay using indirect immunofluorescent antibody staining to search for focus reduction (Higashi, *J. Clin. Microbiol.* 27:2204, 1989, incorporated by reference).

As an alternative to whole cell in vitro assays, purified enzymes isolated from the RRV can be used as targets for rational drug design to determine the effect of the potential drug on enzyme activity, such as thymidylate sunthase or DNA polymerase. The genes for these two enzymes are provided herein. A measure of enzyme activity indicates an effect on the infectious agent itself. Drug screens using herpes viral products are known and have been previously described in EP 0514830 (herpes proteases) and WO 94/04920 (UL 13 gene product).

In particular embodiments, this invention provides an assay for screening anti-KS chemotherapeutics. Infected cells can be incubated in the presence of a chemical agent that is a potential chemotherapeutic against KS (e.g. acyclo-guanosine). The level of virus in the cells is then determined after several days by IFA for antigens or Southern blotting for viral genome or Northern blotting for mRNA and compared to control cells. This assay can quickly screen large numbers of chemical compounds that may be useful against KS. This invention also provides an assay system that is employed to identify drugs or other molecules capable of binding to the DNA molecule or proteins, either in the cytoplasm or in the nucleus, thereby inhibiting or potentiating transcriptional activity. This assay would be useful in the development of drugs that

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are specific against particular cellular activity, or that would potentiate such activity, in time or in level of activity. Also included are drugs identified by this assay which have an anti-viral activity, and an effect against conditions associated with RRV infection, such as KS.

EXAMPLE 24

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Generating Animal Models

Animal models are useful for resolving a number of fundamental problems of infectious diseases that include, but are not limited to, determinants of virulence of the organism, mechanisms of host resistance, mechanisms of pathogenicity, establishment and regulation of chronic infection, and antimicrobial and chemotherapeutic actions of drugs on infectious agents. Variables that are commonly manipulated to address fundamental problems include, but are not limited to, the strain of infectious agent, the infecting dose of infectious agent and the route of administration of the infectious agent, the species or subspecies of animal, the age of animal, and the genetic background of the animal (Viral pathogenesis, N. Nathanson, Lippincot-Raven, Philadelphia, 1997).

In an embodiment in which one or more RRV strains are employed for generating an animal model, the RRV used may be naturally occurring variant isolates recovered from rhesus macaques and other non-human primate species, molecular clones generated from these naturally occurring variant isolates and recombinant viruses with introduced mutations, deletions or recombined genomes designed to address function of specific genes.

By manipulating the infecting dose and route of RRV administration virus-host interactions dependent upon dose and tissue or organ-specific disease manifestations can be explored. Thus, the present invention includes various doses of RRV administered by oral, inhalation, intratracheal, intravaginal, intrarectal and parenteral routes including, but not limited to intravenous, intraarterial, intradermal, subcutaneous, intramuscular, intraperitoneal and organ-specific administration routes such and intracerebral and intraocular administration.

Many disease manifestations with a given infections agent are highly influenced by age and species or subspecies of the host and the particular genetic makeup of the host. The present disclosure provides a virus that is involved in the causation or progression of certain diseases, such as KS, in the rhesus macaque, but is also useful for the study of and discovery of disease manifestations that are host species, age and genetic background dependent. In particular embodiments, one skilled in the art may vary the species of animal to which the RRV is administered to produce or discover a particular disease manifestation, or similarly vary the genetic background of the animal to produce or discover a particular disease manifestation, even including the use of genetically engineered animals.

EXAMPLE 25

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Pharmaceutical Compositions and Modes of Administration

Various delivery systems for administering pharmaceutical proteins from the RRV include encapsulation in liposomes, microparticles, microcapsules, expression by recombinant cells, receptor-mediated endocytosis (see Wu and Wu, *J. Biol. Chem.* 1987, 262:4429-32), and construction of a therapeutic nucleic acid (such as an anti-sense molecule) as part of a retroviral or other vector. Methods of introduction include, but are not limited to, intradermal, intramuscular, intraperitoneal, intravenous, subcutaneous, intranasal, and oral routes. The compounds may be administered by any convenient route, for example by infusion or bolus injection, by absorption through epithelial or mucocutaneous linings (e.g., oral mucosa, rectal and intestinal mucosa, etc.) and may be administered together with other biologically active agents. Administration can be systemic or local. In addition, the pharmaceutical compositions may be introduced into the central nervous system by any suitable route, including intraventricular and intrathecal injection; intraventricular injection may be facilitated by an intraventricular catheter, for example, attached to a reservoir, such as an Ommaya reservoir.

The use of liposomes as a delivery vehicle is another delivery method of the present invention. The liposomes fuse with the target site and deliver the contents of the lumen intracellularly. The liposomes are maintained in contact with the target cells for a sufficient time for fusion to occur, using various means to maintain contact, such as isolation and binding agents. Liposomes may be prepared with purified proteins or peptides that mediate fusion of membranes, such as Sendai virus or influenza virus. The lipids may be any useful combination of known liposome forming lipids, including cationic lipids, such as phosphatidylcholine. Other potential lipids include neutral lipids, such as cholesterol, phosphatidyl serine, phosphatidyl glycerol, and the like. For preparing the liposomes, the procedure described by Kato et al. (*J. Biol. Chem.* 1991, 266:3361) may be used.

The present invention also provides pharmaceutical compositions which include a therapeutically effective amount of one or more RRV proteins or DNA, alone or with a pharmaceutially acceptable carrier.

The pharmaceutical compositions or methods of treatment may be administered in combination with other therapeutic treatments, such as other antineoplastic or antitumorigenic therapies.

Administration of Nucleic Acid Molecules

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In an embodiment in which one or more RRV nucleic acids are employed for generating an animal model, the analog may be delivered intracellularly (e.g., by expression from a nucleic acid vector or by receptor-mediated mechanisms). In a specific embodiment where the therapeutic molecule is a nucleic acid, administration may be achieved by an appropriate nucleic acid expression vector which is administered so that it becomes intracellular, e.g., by use of a retroviral

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vector (see U.S. Patent No. 4,980,286), or by direct injection, or by use of microparticle bombardment (e.g., a gene gun; Biolistic, Dupont), or coating with lipids or cell-surface receptors or transfecting agents, or by administering it in linkage to a homeobox-like peptide which is known to enter the nucleus (see e.g., Joliot et al., *Proc. Natl. Acad. Sci. USA* 1991, 88:1864-8). Alternatively, the nucleic acid can be introduced intracellularly and incorporated within host cell DNA for expression, by homologous recombination.

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The vector pCDNA, is an example of a method of introducing the foreign cDNA into a cell under the control of a strong viral promoter (CMV) to drive the expression. However, other vectors can be used. Other retroviral vectors (such as pRETRO-ON, Clontech), also use this promoter but have the advantages of entering cells without any transfection aid, integrating into the genome of target cells ONLY when the target cell is dividing (as cancer cells do, especially during first remissions after chemotherapy) and they are regulated. It is also possible to turn on the expression of the RRV nucleic acid by administering tetracycline when these plasmids are used. Hence these plasmids can be allowed to transfect the cells, then administer a course of tetracycline with a course of chemotherapy to achieve better cytotoxicity.

Other plasmid vectors, such as pMAM-neo (also from Clontech) or pMSG (Pharmacia) use the MMTV-LTR promoter (which can be regulated with steroids) or the SV10 late promoter (pSVL, Pharmacia) or metallothionein - responsive promoter (pBPV, Pharmacia) and other viral vectors, including retroviruses. Examples of other viral vectors include adenovirus, AAV (adenoassociated virus), recombinant HSV, poxviruses (vaccinia) and recombinant lentivirus (such as HIV). All these vectors achieve the basic goal of delivering into the target cell the cDNA sequence and control elements needed for transcription. The present invention includes all forms of nucleic acid delivery, including synthetic oligos, naked DNA, plasmid and viral, integrated into the genome or not.

Also contemplated are inhibitory nucleic acid therapeutics which can inhibit the activity of RRV, for example in subject with KS or other diseases associated with RRV infection. Inhibitory nucleic acids may be single-stranded nucleic acids, which can specifically bind to a complementary nucleic acid sequence. By binding to the appropriate target sequence, an RNA-RNA, a DNA-DNA, or RNA-DNA duplex or triplex is formed. These nucleic acids are often termed "antisense" because they are usually complementary to the sense or coding strand of the gene, although recently approaches for use of "sense" nucleic acids have also been developed. The term "inhibitory nucleic acids" as used herein, refers to both "sense" and "antisense" nucleic acids.

By binding to the target nucleic acid, the inhibitory nucleic acid can inhibit the function of the target nucleic acid. This could, for example, be a result of blocking DNA transcription, processing or poly(A) addition to mRNA, DNA replication, translation, or promoting inhibitory mechanisms of the cells, such as promoting RNA degradation. Inhibitory nucleic acid methods therefore encompass a number of different approaches to altering expression of RRV genes.

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Cleavage, and therefore inactivation, of the target nucleic acids may be effected by attaching a substituent to the inhibitory nucleic acid which can be activated to induce cleavage reactions. The substituent can be one that affects either chemical, or enzymatic cleavage. Alternatively, cleavage can be induced by the use of ribozymes or catalytic RNA. In this approach, the inhibitory nucleic acids would include either naturally occurring RNA (ribozymes) or synthetic nucleic acids with catalytic activity.

The inhibitory nucleic acid therapies can be used to target nucleic acids to sequences of RRV for use in treating conditions caused by the RRV, or proteins of the RRV, for example for treating KS or KS-like syndromes.

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Administration of Antibodies

Therapeutic, intravenous, polyclonal or monoclonal antibodies has been used as a mode of passive immunotherapy of herpesviral diseases, such as infection with CMV. Immune globulin from subjects previously infected with the RRV and bearing a suitably high titer of antibodies against the virus can be given in combination with antiviral agents (e.g. ganciclovir), or in combination with other modes of immunotherapy that are currently being evaluated for the treatment of KS, which are targeted to modulating the immune response (i.e. treatment with copolymer-1, antiidiotypic monoclonal antibodies, T cell "vaccination"). Antibodies specific for an epitope expressed on cells infected with the RRV are preferred and can be obtained as described above.

The present invention also provides pharmaceutical compositions which include a therapeutically effective amount of the antibody, and a pharmaceutically acceptable carrier or excipient.

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EXAMPLE 26

Vaccines

This invention provides substances suitable for use as vaccines for the prevention of diseases associated with RRV infection, such as KS, and methods for administering them. The vaccines are directed against RRV, and may include antigens obtained from RRV. In one embodiment, the vaccine contains attenuated RRV. In another embodiment, the vaccine contains killed RRV. In another embodiment, the vaccine contains a nucleic acid vector encoding RRV, or a surface protein, such as a capsid protein. In another embodiment, the vaccine is a subunit vaccine containing an RRV subunit, such as glycoprotein B, major capsid protein, or other gene products found to elicit appropriate humoral and/or cell mediated immune responses.

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This invention also provides a method of vaccinating a subject against Kaposi's sarcoma and lymphoproliferative disorders, comprising administering to a susceptible subject an effective amount of the peptide or polypeptide encoded by an isolated DNA molecule encoding a

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polypeptide or combination of polypeptides expressed by the DNA molecule, and a suitable acceptable carrier. In one embodiment, naked DNA is administered to the subject in an effective amount to vaccinate the subject against Kaposi's sarcoma and lymphoproliferative disorders, or other disease associated with RRV infection.

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The vaccine can be made using synthetic peptide or recombinantly-produced polypeptide described above as antigen. Typically, a vaccine will include from about 1 to 50 micrograms of antigen, for example from about 15 to about 45 micrograms. Typically, the vaccine is formulated so that a dose includes about 0.5 milliliters. The vaccine may be administered by any route known in the art, for example parenteral, subcutaneous or intramuscular.

There are a number of strategies for amplifying an antigen's effectiveness, particularly as related to the art of vaccines. For example, cyclization of a peptide can increase the peptide's antigenic and immunogenic potency. See U.S. Pat. No. 5,001,049. More conventionally, an antigen can be conjugated to a suitable carrier, usually a protein molecule. This procedure can allow multiple copies of an antigen, such as a peptide, to be conjugated to a single larger carrier molecule. Additionally, the carrier may possess properties which facilitate transport, binding, absorption or transfer of the antigen.

For parenteral administration, such as subcutaneous injection, examples of suitable carriers are the tetanus toxoid, the diphtheria toxoid, serum albumin and lamprey, or keyhole limpet, hemocyanin because they provide the resultant conjugate with minimum genetic restriction. Conjugates including these universal carriers can function as T cell clone activators in individuals having very different gene sets. The conjugation between a peptide and a carrier can be accomplished using one of the methods known in the art. Specifically, the conjugation can use bifunctional cross-linkers as binding agents as detailed, for example, by Means and Feeney, "A recent review of protein modification techniques," *Bioconjugate Chem.* 1:2-12 (1990).

Vaccines against RRV can be made from the RRV envelope glycoproteins. These proteins can be purified and used for vaccination (Lasky, L. A., 1990, *J. Med. Virol.* 31:59). MHC-binding peptides from cells infected with the human herpesvirus can be identified for vaccine candidates per the methodology of Marloes, et al., 1991, *Eur. J. Immunol.* 21:2963-2970. The RRV antigen may be combined or mixed with various solutions and other compounds as is known in the art. For example, it may be administered in water, saline or buffered vehicles with or without various adjuvants or immunodiluting agents. Examples of such adjuvants or agents include aluminum hydroxide, aluminum phosphate, aluminum potassium sulfate (alum), beryllium sulfate, silica, kaolin, carbon, water-in-oil emulsions, oil-in-water emulsions, muramyl dipeptide, bacterial endotoxin, lipid X, Corynebacterium parvum (Propionibacterium acnes), Bordetella pertussis, polyribonucleotides, sodium alginate, lanolin, lysolecithin, vitamin A, saponin, liposomes, levamisole, DEAE-dextran, blocked copolymers or other synthetic adjuvants. Such adjuvants are available commercially from various sources, for example, Merck Adjuvant 65

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(Merck and Company, Inc., Rahway, N.J.) or Freund's Incomplete Adjuvant and Complete Adjuvant (Difco Laboratories, Detroit, Mich.). Other suitable adjuvants are Amphigen (oil-inwater), Alhydrogel (aluminum hydroxide), or a mixture of Amphigen and Alhydrogel. Only aluminum is approved for human use.

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The proportion of antigen and adjuvant can be varied over a broad range so long as both are present in effective amounts. For example, aluminum hydroxide can be present in an amount of about 0.5% of the vaccine mixture (AhO3 basis). On a per-dose basis, the amount of the antigen can range from about 0.1 µg to about 100 µg protein per subject, for example about 1 µg to about 50 µg per dose, or about 15 µg to about 45 µg. A suitable dose size is about 0.5 ml. Accordingly, a dose for intramuscular injection, for example, would comprise 0.5 ml containing 45 µg of antigen in admixture with 0.5% aluminum hydroxide. After formulation, the vaccine may be incorporated into a sterile container which is then sealed and stored at a low temperature, for example 4°C., or it may be freeze-dried. Lyophilization permits long-term storage in a stabilized form.

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The vaccines may be administered by any conventional method for the administration of vaccines including oral and parenteral (e.g., subcutaneous or intramuscular) injection.

Intramuscular administration is preferred. The treatment may consist of a single dose of vaccine or a plurality of doses over a period of time. Also, the antigen could be a component of a recombinant vaccine which could be adaptable for oral administration. Vaccines of the invention may be combined with other vaccines for other diseases to produce multivalent vaccines. A pharmaceutically effective amount of the antigen can be employed with a pharmaceutically acceptable carrier such as a protein or diluent useful for the vaccination of mammals, particularly humans. Other vaccines may be prepared according to methods well-known to those skilled in the art.

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Those of skill will readily recognize that it is only necessary to expose a mammal to appropriate epitopes in order to elicit effective immunoprotection. The epitopes are typically segments of amino acids which are a small portion of the whole protein. Using recombinant genetics, it is routine to alter a natural protein's primary structure to create derivatives embracing epitopes that are identical to or substantially the same as (immunologically equivalent to) the naturally occurring epitopes. Such derivatives may include peptide fragments, amino acid substitutions, amino acid deletions and amino acid additions of the amino acid sequence for the viral polypeptides from the human herpesvirus. For example, it is known in the protein art that certain amino acid residues can be substituted with amino acids of similar size and polarity without an undue effect upon the biological activity of the protein. The human herpesvirus proteins have significant tertiary structure and the epitopes are usually conformational. Thus, modifications should generally preserve conformation to produce a protective immune response.

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EXAMPLE 27

Peptide Synthesis and Purification

The peptides provided by the present invention can be chemically synthesized by any of a number of manual or automated methods of synthesis known in the art. For example, solid phase peptide synthesis (SPPS) is carried out on a 0.25 millimole (mmole) scale using an Applied Biosystems Model 431A Peptide Synthesizer and using 9-fluorenylmethyloxycarbonyl (Fmoc) amino-terminus protection, coupling with dicyclohexylcarbodiimide/ hydroxybenzotriazole or 2-(1H-benzo-triazol-1-yl)-1,1,3,3-tetramethyluronium hexafluorophosphate/ hydroxybenzotriazole (HBTU/HOBT), and using p-hydroxymethylphenoxymethylpolystyrene (HMP) or Sasrin resin for carboxyl-terminus acids or Rink amide resin for carboxyl-terminus amides.

Fmoc-derivatized amino acids are prepared from the appropriate precursor amino acids by tritylation and triphenylmethanol in trifluoroacetic acid, followed by Fmoc derivitization as described by Atherton et al. (Solid Phase Peptide Synthesis, IRL Press: Oxford, 1989).

Sasrin resin-bound peptides are cleaved using a solution of 1% TFA in dichloromethane to yield the protected peptide. Where appropriate, protected peptide precursors are cyclized between the amino- and carboxyl-termini by reaction of the amino-terminal free amine and carboxyl-terminal free acid using diphenylphosphorylazide in nascent peptides wherein the amino acid sidechains are protected.

HMP or Rink amide resin-bound products are routinely cleaved and protected sidechain-containing cyclized peptides deprotected using a solution comprised of trifluoroacetic acid (TFA), optionally also comprising water, thioanisole, and ethanedithiol, in ratios of 100:5:5:2.5, for 0.5-3 hours at room temperature.

Crude peptides are purified by preparative high pressure liquid chromatography (HPLC), for example using a Waters Delta-Pak C18 column and gradient elution with 0.1% TFA in water modified with acetonitrile. After column elution, acetonitrile is evaporated from the eluted fractions, which are then lyophilized. The identity of each product so produced and purified may be confirmed by fast atom bombardment mass spectroscopy (FABMS) or electrospray mass spectroscopy (ESMS).

Having illustrated and described the principles of cloning the RRV genome, cDNA, proteins encoded by the cDNA, and modes of use of these biological molecules, it should be apparent to one skilled in the art that the invention can be modified in arrangement and detail without departing from such principles. In view of the many possible embodiments to which the principles of our invention may be applied, it should be recognized that the illustrated embodiments are only examples of the invention and should not be taken as a limitation on the scope of the invention. Rather, the scope of the invention is in accord with the following claims. We therefore claim as our invention all that comes within the scope and spirit of these claims.

CLAIMS AS AMENDED UNDER ARTICLE 34

- 5 1. An isolated virus (RRV) as deposited with ATCC as deposit accession number VR-2601.
 - 2. A purified virus, having a nucleic acid sequence
 - (a) shown in SEQ ID NO 1 or
 - (b) a conservative variant thereof.
 - 3. The purified virus of claim 2, wherein the nucleic acid sequence has at least 95% sequence identity to the nucleic acid sequence shown in SEQ ID NO 1.
 - 4. A purified protein encoded by an open reading frame of the virus of claim 2.
 - 5. A purified protein of claim 4, wherein the protein comprises an amino acid sequence selected from the group consisting of:
 - (a) an amino acid sequence shown in odd numbered sequences of SEQ ID NOS. 3-165; and
 - (b) amino acid sequences that differ from those specified in (a) by one or more conservative amino acid substitutions wherein the function of the protein is preserved.
 - 6. A purified protein with an amino acid sequence that is at least 95% sequence identity to the sequences specified in claim 5(a) or 5(b).
- 7. The purified protein of claim 5, wherein the amino acid sequence is selected from odd numbered sequences within the group consisting of SEQ ID NOS 3-19 and 23-165.
 - 8. An isolated nucleic acid molecule encoding a protein according to claim 5.
 - 9. An isolated nucleic acid molecule according to claim 8, wherein the molecule comprises a sequence selected from the group consisting of even numbered sequences of SEQ ID NOS 2-164.

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- 10. The isolated nucleic acid molecule according to claim 9, wherein the molecule comprises a sequence selected from the group consisting of even numbered sequences of SEQ ID NOS 2-18 and 22-164.
- 11. A recombinant nucleic acid molecule comprising a promoter sequence operably linked to a nucleic acid molecule according to claim 8.
- 12. A cell transformed with a recombinant nucleic acid molecule according to claim 8.
 - 13. A non-human mammal purposefully infected with the virus of claim 2.
 - 14. The mammal of claim 13, wherein the mammal is a primate.
- - (a) at least 20 contiguous nucleotides of the nucleic acid sequence of the virus of claim 2;
 - (b) at least 30 contiguous nucleotides of the nucleic acid sequence of the virus of claim 2; and
 - (c) at least 50 contiguous nucleotides of the nucleic acid sequence of the virus of claim 2.
 - 16. An isolated nucleic acid molecule that:
 - (a) hybridizes under stringent conditions with a nucleic acid probe comprising the sequence of claim 15; and
 - (b) encodes a protein of claim 6.
 - 17. An isolated nucleic acid molecule encoding a protein of claim 6.
 - 18. An isolated nucleic acid molecule encoding all proteins encoded by the virus of claim 2, and having a biological activity of an RRV virus.
- 19. A method for testing the efficacy of a drug in the treatment of a condition associated with the virus of claim 2, the method comprising:
 - (a) administering the drug to a non-human primate infected with the virus of claim 2; and

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(b) observing the primate to determine if the drug prevents or reduces the presentation of one or more symptoms associated with viral infection.

- 20. The method of claim 19, wherein the primate is immunocompromised.
- 21. The method of claim 20, wherein the drug is for the treatment of Kaposi's sarcoma and lymphoproliferative disorders.
- The method of claim 20, wherein the primate is immuno-compromised as a result of infection by Simian Immunodeficiency Virus (SIV).
 - 23. The method of claim 19, wherein the condition associated with infection with the virus of claim 2 is one or more of B-cell hyperplasia, lymphadenopathy, splenomegaly, hypergammaglobinulinemia or autoimmune hemolytic anemia.
 - 24. The method of claim 19, wherein the non-human primate is a Rhesus macaque monkey.
 - 25. A method for producing a non-human primate model for testing potential treatments for a condition associated an infection with the virus of claim 2, comprising

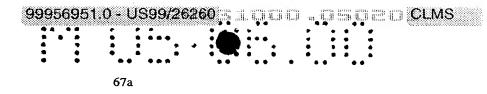
 (a) administering a treatment to the primate to render the primate immunocompromised; and
 - (b) infecting the primate with the virus of claim 2.
- 25 26. The method of claim 25, wherein the condition is Kaposi's sarcoma and lymphoproliferative disorders.
 - 27. The method of claim 25 wherein the treatment used to render the primate immuno-compromised is infection with SIV.
 - 28. The method of claim 25 wherein the non-human primate is a Rhesus macaque monkey.
- 29. A method for testing the efficacy of a candidate vaccine against the virus of claim 2, or conditions associated infection with virus of claim 2, the method comprising:

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- (a) administering the vaccine to a subject capable of infection with the virus of claim 2;
 - (b) inoculating the subject with the virus; and
- (c) observing the subject to determine if the vaccine prevents or reduces an incidence of viral infection or presentation of one or more conditions associated with the viral infection.
 - 30. The method of claim 29, wherein the subject is a primate.
 - 31. The method of claim 30, wherein the primate is a non-human primate.
 - 32. The method of claim 29, wherein the primate is immunocompromised.
- The method of claim 29, wherein the conditions associated with infection include B-cell hyperplasia, lymphadenopathy, splenomegaly, hypergammaglobinulinemia or autoimmune hemolytic anemia.
 - 34. The method of claim 31, wherein the non-human primate is a Rhesus macaque monkey.

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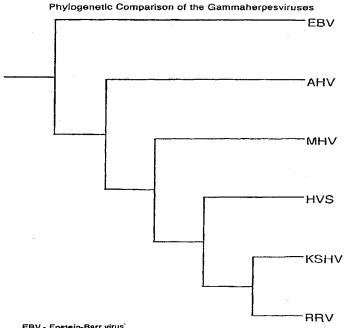
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(54) Title: CLONING OF RHADINOVIRUS GENOME AND METHODS FOR ITS USE

(57) Abstract

A novel rhesus macaque rhadinovirus, herein designated RRV, is disclosed. The genomic, cDNA and proteins sequences are provided. RRV has some similarity to human Kaposi's sarcoma-associated herpesvirus and causes Kaposi's sarcoma-like symptoms in immuno-compromised non-human primates. RRV possesses genes for both Interleukin-6 and macrophage inflammatory protein 1. The genome of RRV is useful for research, clinical and diagnostic applications aimed towards the rhadinoviruses and herpesviruses in general and KSHV in particular. In addition, methods for using RRV to produce a non-human primate model for the testing of Kaposi's sarcoma-associated herpesvirus therapeutics and vaccines are presented.



EBV - Epstein-Barr virus'

AHV - Alcelaphine herpesvirus

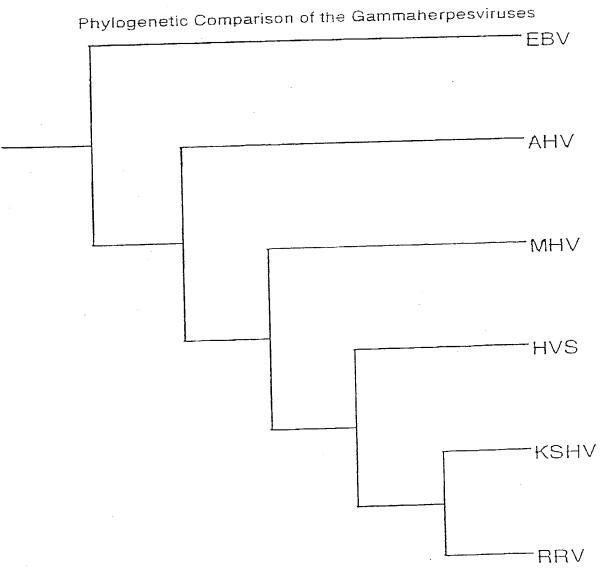
MHV - Murine herpesvirus 68 HVS - Herpesvirus saimiri

KSHV - Kaposi's sarcoma-associated herpesvirus

RRV - Rhesus rhadinovirus 17577

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FIG. 1



EBV - Epstein-Barr virus

AHV - Alcelaphine herpesvirus

MHV - Murine herpesvirus 68

HVS - Herpesvirus saimiri

KSHV - Kaposi's sarcoma-associated herpesvirus

RRV - Rhesus rhadinovirus 17577

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FIG. 2

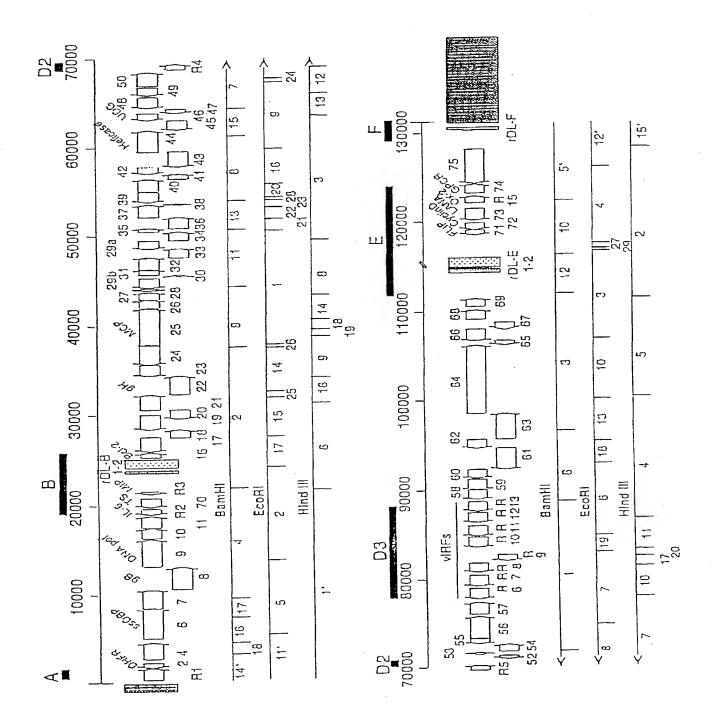
Restriction Fragments of the RRV 17577 Genome

Bar	пНI	Eco	oRI .		Hine	ત્રાાા દ
fragment number	fragment size (bp)	fragment number	fragment size (bp)	,	fragment number	fragment size (bp)
1 2 3 4 5* 6 7 8 9 10 11 12 13 14* 15 16 17 18	17189 15598 15441 12360 8943 7747 7718 7142 6667 6474 6333 3978 3411 3157 3008 2916 2210 1343	1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29	12476 10342 9565 9213 8465 8036 7969 7416 7278 7002 5400 5054 4907 4771 4272 4099 3516 2102 1868 1603 1512 1221 910 624 609 592 584 122 107		1° 2 3 4 5 6 7 8 9 10 11 12 13 14° 15 16 17 18 19	22006 17108 16542 14134 11516 10743 8452 5995 4679 3374 2963 2891 2849 2832 1599 1272 1016 853 811

^{*} Indicates that the fragment size excludes terminal repeat sequences

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FIG. 3



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FIG. 4

MacVector Output for long unique region of rhesus rhadinovirus 17577

LOCUS &	LONG	UNIQU	131634	BP	P DS-INA UPDATED 06/26/98
DEFINITION					
ACCESSION	-				. *
REYWORDS	_				
SOURCE	-				
FEATURES					Description
Jcpq .				1	R1
bebt				(CI	Similar to HHV8 Orf 2 - dihydrofolate reductase
pept				1	Similar to HAV8 Orf 4 - complement binding
					protein
pept				1	Similar to HHV8 Orf 6 - ssDNA binding protein
pept				1	Similar to HWW8 Orf 7 - transport protein
pept				1	Similar to HHV8 Orf 8 - glycoprotein 3
pept					Similar to HHV8 Orf 9 - DEA polymerase
pept					Similar to HHVB Orf 10
acac					Similar to HHV8 Orf 11
pept					1 P2 viral II-6
ည်ဧညင				(CI	Similar to SHV8 Orf 70
pept				(C]	R3 similar to HAV8 MIP misc. feature MIP homology, but no initiation
īrag					
					cocon
عفد					xepeat sequence
= p€				_	repeat sequence I Similar to HRV8 Orf 15 - Bol-2 homolog
pept				1	Similar to MEV8 Out 17 - capsid protein
pept		•			Similar to mive Orf 18
pept				۲.	l Similar to mive Orf 19 - tegiment protein
pept					Similar to miv Ori 20
ည်ဧညင				(C1	Similar to MAVS Orf 21 - thymidine kinese
హేకఫ్ర				7	1 Similar to HRV8 Orf 22 - glycoprotein H
bebt				/~1	Similar to HTV8 Orf 23
				(C1	I Similar to HAV8 Orf 24
pept				1	Similar to HEV8 Orf 25 - major capsid protein
pept				1	Similar to HEV8 Orf 25 - capsid protein
pept					Similar to HAVE Orf 27
pept					Similar to HHV8 Orf 28
pept					Similar to HAV8 Orf 295
pept pept					Similar to HAV8 Orf 30
					Similar to HEV8 Orf 31
pept pept					l Similar to Have Orf 32
ခံခေင ခဲ့ခေင					1 Similar to HW8 Orf 33
pept					1 Similar to HTV8 Orf 29a
pept				1	1 Similar to HIV8 Orf 34
ಕ್ಕಾರ				1	l Similar to HHV8 Orf.35
pept				1	l Similar to HTV8 Orf 36 -kinase
pept				1	1 Similar to HAV8 Orf 37 - alkeline exomuclease
pept				.1	1 Similar to Have Orf 38
pept				(C1	1 Similar to HAV8 Orf 39 - glycoprotein M
pept				1	1 Similar to HEV8 Orf 40 - helicase - primese
pept				1	l Similar to MAVB Orf il - helicese - primase
pept				(C1	1 Similar to HAVE Or! 42
. pept				(C1	1 Similar to HTV8 Orf 43 - capsid protein
pept				1	1 Similar to serva Orf 44 - helicese -primase
pept				(Cl	1 Similar to MTV3 Orf 45

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FIG. 5

Comparison of Corresponding Repeats in RRV and KSHV

virus	insert name	total length	repeat unit length	G + C content
KSHV	frnk¹	332 bp 292 bp	20 bp 30 bp	80.1% 84.9%
RRV	syko¹	304 bp 1008 bp	26 bp 25 bp	53.3% 79.9%
KSHV	zppa¹	308 bp 244 bp	23 bp 23 bp	74.0% 77.9%
RRV	vrtgo ¹	405 bp 1029 bp	19 bp 32 bp	74.6% 84.4%
virus	insert name	total length	repeat unit length	G + A content
KSHV	mdsk	409 bp	2	75.4%
RRV	brds	196	13 bp	81.6%

¹ KSHV fmk and zppa and RRV syko and vrtgo are tandem repeats.

² KSHV mdsk is a complex repeat with no defined unit length.

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FIG. 6

	Rh R13								28.818	58.103 52.964	1	31.124 25.072		100,000	
	Rh R12							33.038	61.254 50.997	30,364 18.623	33.526 25.723	33.923 23.849	100.000		
	Rh R11	29.972 21.849		1				50.773	35.693 23.849	28.216 21.577	32.951 23.496	100.000			
KSHVa	Rh R10	33.705 · 26.184					54.427 47.917		31.412 24.207	28,980 18,367	100.000 100.000				
RRV and	Rh R9						29.918 22,131		,	100.000					
oded by	Rh RB	28.857 19,427				**	26.393	34.513 26.254	100.000						
ements c	Rh R7	28.291 20.728						100.000							
ulatory el	Rh RG	25.044 21.130					100.000								
on of Interferon regulatory elements coded by RRV and KSHVa	KSHV K11				32.036 21.895	100.000 100.000									
n of Inter	KSHV K10.5				100,000										
Comparisor	KSHV K10.1			100.000											
	KSHV K10		100.000												
	KSHV K9	100.000 100.000													
1			KSHV K9	KSHV K10	KSHV K10.1	ភ	KSHV K11	Rh R6					Kh Kan		Rh R13

a. Blank cells Indicated no similarity; upper number is percent similarity; lower number is percent Identity.

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										,	<i>/</i> I	,											ĺ.							
		Putative Function		Dihydrofolate reductase Complement binding protein	:	ssDNA binding protein	Transport protein	Glycoprotein B	DNA polymerase				Thymidylate synthase		Bcl-2 homolog	Capsid protein		Tegument protein	:	Thymidine kinase	Glycoprotein H			Major capsid protein	Capsid protein			Packaging protein		
		% –		54.8% 35.3%	38.6%	53.5%	47.7%	53.1%	62.5%	23.3%	32.4%		64.6%		21.4%	42.2%	48.8%	46.9%	35.6%	31.7%	31.5%	29.8%	46.8%	67.5%	58.2%	27.1%		62.9%	29.2%	39.9%
Fs	HVS	% w		65.6% 42.0%	44.0%	65.2%	58.1%	62.4%					72.1%		31.4%	49.0%	60.2%	55.5%	43.2%	39.0%	42.3%	40.5%	56.3%	%2'92	69.1%	35.0%		74.4%	40.3%	20.5%
A V and HVS OR		Size aa		187 360	287	1128	629	808	1009	407	405		294		160	475	256	543	303	527	717	253	731	1371	304	280	93	387	75	208
FIG. 7A Comparison of RRV, KSHV and HVS ORFs		% I		46.0% 35.7%		63.3%	51.5%	65.5%	%0.79	34.8%	31.7%		66.1%	32.3%	46.0%	44.3%	28.0%	52.8%	44.7%	44.6%	40.7%	48.5%	58.7%	72.5%	64.3%	25.3%	26.5%	66.4%	38.2%	45.4%
Сотраг	KSHV	Sim		55.1% 40.9%		71.3%	60.1%	73.3%	75.0%	43.5%	41.3%		72.1%	41.9%	28.0%	20.6%	68.1%	61.1%	51.8%	54.0%	50.1%	26.8%	86.3%	79.9%	71.8%	33.6%	30.1%	%9′.22	51.3%	%0.99
		Size		210	 - 	1133	695	845	1312	418	407		337	92	175	553	257	549	320	280	730	404	752	1376	305	290	102	351	77	224
		Size aa	423	188 645	<u>!</u>	1132	989	829	1014	384	409	207	333	115	187	536	299	547	350	557	704	402	732	1378	307	269	91	348	92	217
		Strand	+	. +		+	+	+	+	+	+	•	•	•	+		+	•		+	+		,	+	+	+	+	•	+	+
		ORF	R1	Orf 2	<u> </u>	Orf 6	Orf 7	Orf 8	Orf 9	Orf 10	Orf 11	R21	Orf 70	R33	Orf 16	Orf 17	Orf 18	Orf 19	Orf 20	Orf 21	Orf 22	Orf 23	Orf 24	Orf 25	Orf 26	Orf 27	Orf 28	Orf 29b	Orf 30	Orf 31

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												8.	/1	3																		
	Putative Function			Packaging protein			Kinase	Alkaline exonuclease		Glycoprotein M	Helicase-primase	Helicase-primase		Capsid protein	Helicase-primase		Uracil DNA glucosidase	Glycoprotein L			Transactivator					dUTPase		DNA replication protein	Immediate-early protein			
	% -	34.1%	39.1%	49.8%	40.6%	37.4%	28.7%	53.2%	34.8%	27.0%	28.1%	29.1%	38.1%	%9.99	62.6%		59.1%	23.9%	25.8%	23.3%	21.6%			30.4%	28.9%	36.4%	44.4%	43.6%	31.5%			
HVS	% w	43.2%	49.1%	57.8%	53.7%	51.0%	38.4%	63.0%	39.4%	67.1%	39.1%	37.3%	51.2%	66.4%	71.1%		67.5%	33.3%	34.1%	35.1%	29.7%			41.7%	43.3%	46.5%	52.5%	54.0%	40.3%			
	Size aa	441	330	303	316	150	431	483	99	366	420	161	265	563	781	257	252	141	797	303	535			115	06	287	200	835	416			
	% –	41.8%	42.1%	61.2%	48.5%	35.6%	46.1%	63.5%	45.0%	59.3%	32.7%	26.0%	46.1%	61.6%	%0.99	24.9%	60.1%	27.7%	29.2%	54.2%	37.8%			45.4%	46.2%	41.0%	55.2%	52.5%	47.1%	21.1%	20.7%	19.4%
KSHV	Sim %	49.9%	52.1%	%2.99	28.9%	47.7%	26.0%	72.4%	26.7%	73.0%	42.2%	33.5%	56.8%	69.7%	73.9%	31.2%	71.9%	31.9%	36.2%	66.1%	46.6%			58.5%	51.0%	48.6%	62.9%	61.2%	%9.09	26.0%	28.3%	28.9%
	Size aa	454	312	312	327	151	444	486	61	399	457	205	278	605	788	407	255	167	402	302	631			131	110	318	227	843	275			
	Size aa	464	336	327	327	149	435	480	69	378	468	203	272	576	790	352	255	169	389	301	514	206	111	139	104	290	210	828	442	415	415	351
RRV	Strand	+	+		+	+	+	+	+	•	+	+	•	ı	+	1	•	1	1		+	+	+		•	+		+	+	•	:	•
	ORF	Orf 32	Orf 33)rf 29a	Orf 34	Orf 35	Orf 36	Orf 37	Orf 38	Orf 39	Orf 40	Orf 41	Orf 42	Orf 43	Orf 44	Orf 45	Orf 46	Orf 47	Orf 48	Orf 49	Orf 50	R41	R51	Orf 52	Orf 53	Orf 54	Orf 55	Orf 56	Orf 57	R64	R74	R84

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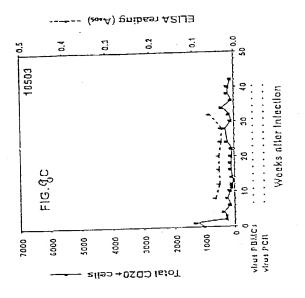
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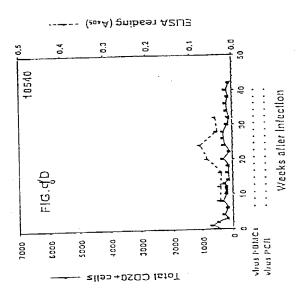
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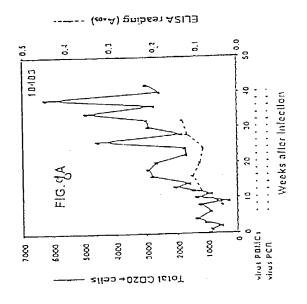
										9/	1	3.														
	Putative Function							DNA replication protein	Ribonucleotide reductase,	small small small small	Niboliucieoliue Teduciase,	large Assembly / DNA maturation	Tegument protein	Tegument protein	Capsid protein	•	Tegument protein	Glycoprotein			Flip homolog	Cyclin D homolog	Immediate-early gene		G-protein coupled receptor	Tegument protein / FGARAT
	% –						29.5%	32.7%	62.4%	F3 30/	02.570	41.9%	34.6%	29.4%	33.1%	32.3%	51.4%	44.3%	49.0%		15.1%	29.2%	20.8%		32.1%	34.4%
HVS	% v						39.9%	40.7%	71.0%	64 40/	04.4%	53.8%	43.4%	39.2%	41.0%	43.6%	28.6%	53.5%	57.5%		25.3%	37.5%	29.0%		41.1%	43.2%
	Size aa						357	368	302	787	/0/	330	833	2469	139	435	253	436	261		167	254	407		321	1299
	% –		26.2%	21.8%			38.2%	51.8%	%0.02	61 70/	%/:10	56.5%	42.6%	40.2%	38.6%	46.4%	64.7%	44.8%	65.5%		30.9%	38.6%	16.8%	31.2%	41.1%	44.0%
KSHV	% Sim		33.7%	30.0%			45.2%	60.3%	78.2%	/ac 03	03.5%	64.4%	51.8%	49.6%	48.2%	51.9%	%9.69	53.2%	73.1%		38.8%	49.8%	23.6%	35.2%	51.6%	52.2%
	Size aa						357	396	302	402	767	334	927	2635	170	429	271	545	225		139	257	1162	348	342	1296
	Size aa	253	385	390	355	364	360	394	314	700	/88	331	939	2548	169	448	224	457	297	228	174	254	447	253	342	1298
RRV	Strand		ı	•	,		•	•					+	+		,	•	+	+			•	ı	+	+	,
	ORF	R94	R104	R114	R124	R134	Orf 58	Orf 59	Orf 60	3	0H 61	Orf 62	Orf 63	Orf 64	Orf 65	Orf 66	Orf 67	Orf 68	Orf 69	R141	Orf 71	Orf 72	Orf 73	R155	Orf 74	Orf 75

% Sim., percent similar; % ld., percent identical; ssDNA, single-stranded DNA; FGARAT, N-formalglycinamide ribotide amidotransferase; 1, no similarity found; 2, compared to HVS ORF 4a and 4b; 3, compared to KSHV K9; 5, compared to KSHV K14.

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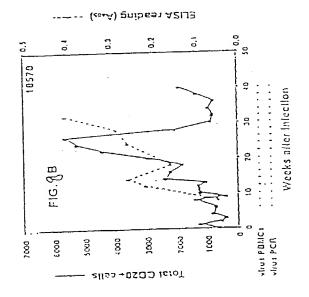


FIG. 8

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18483
18503
18540
18570

10-

CD4+

	PE	BLs			LNI	ЛCs	
18483	18503	18540	18570	18483	18503	10540	18570

CD8+

CD20÷

RhKSHV MIP

0

β-globin



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.

FIG. 10

atg Met 1	ttc Phe	cct Pro	gtc Val	tgg Trp 5	ttc Phe	gtc Val	ttg Leu	ttt Phe	tac Tyr 10	ctg Leu	tcg Ser	tgt Cys	tgg Trp	gcg Ala 15	gcc Ala	48
agc Ser	cct Pro	acg Thr	ctg Leu 20	gcg Ala	cct Pro	ccc	ccg Pro	act Thr 25	gcc Ala	gct Ala	gga Gly	att Ile	aac Asn 30	gtt Val	ctc Leu	96
ccc Pro	cag Gln	tgg Trp 35	gcc Ala	ggc	aac Asn	cgc Arg	gcc Ala 40	tct Ser	ctt Leu	gac Asp	agg Arg	acc Thr 45	agg Arg	Gly 999	cgc Arg	144
ctg Leu	tct Ser 50	gaa Glu	gtg Val	ejy aaa	tta Leu	aac Asn 55	ata Ile	cag Gln	cgc Arg	tgg Trp	ttc Phe 60	gtt Val	tac Tyr	ctg Leu	tgc Cys	192
cac His 65	cac His	tcc Ser	act Thr	ctc Leu	tgt Cys 70	cgg Arg	gtg Val	cgt Arg	gag Glu	tac Tyr 75	ccg Pro	cgc Arg	atc Ile	atg Met	tcg Ser 80	240
ttt Phe	gtt Val	cac His	ttc Phe	cct Pro 85	ata Ile	ttg Leu	atg Met	tct Ser	aac Asn 90	gtt Val	gag Glu	tgc Cys	cag Gln	cgc Arg 95	Arg cgc	288
gag Glu	ttt Phe	cgc Arg	100 GJA 333	gcc Ala	gag Glu	tgt Cys	atg Met	aac Asn 105	Ala	atg Met	gtt Val	cgc Arg	999 Gly 110	cțc Leu	cgg Arg	336
					ctg Leu											384
ccc Pro	999 Gly 130	gac Asp	gcg Ala	gac Asp	gcc Ala	gcg Ala 135	gcc Ala	att Ile	ggc Gly	tcc Ser	gcg Ala 140	gtg Val	acc Thr	gtg Val	gtg Val	432
ctg Leu 145	tcc Ser	gcc Ala	ctc Leu	gac Asp	tct Ser 150	cta Leu	att Ile	gag Glu	gag Glu	ctt Leu 155	ccc Pro	gta Val	aat Asn	aac Asn	aag Lys 160	480
ata Ile	ggt Gly	Gly	gcg Ala	gag Glu 165	tct Ser	aat Asn	gaa Glu	aaa Lys	acc Thr 170	gtg Val	cgt Arg	gcg Ala	ttg Leu	gga Gly 175	cj aaa	528
cag Gln	agc Ser	ccc Pro	cgg Arg 180	gac	gtt Val	gtt Val	ctc Leu	agc Ser 185	gcg Ala	ttt Phe	cgc Arg	ata Ile	ctg Leu 190	gaa Glu	tat Tyr	576
cta Leu	cag Gln	atg Met 195	ttt Phe	ttg Leu	cgg Arg	gac Asp	999 Gly 200	cgc Arg	cgc Arg	gca Ala	ata Ilc	gct Ala 205	atg Met	atg Met	taa	624

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FIG. 11

											gtg Val					₫ 8
											gga Gly					96
ctc Leu	tgc Cys	tgt Cys 35	ttg Leu	61A a aa	tat Tyr	gta Val	act Thr 40	cat His	ctg Leu	ccg Pro	cca Pro	ccc Pro 45	ggt Gly	tta Leu	gtg Val	144
											gtg Val 60					192
											aat Asn					240
											cgt Arg					288
aga Arg	aga Arg	acc Thr	cgg Arg 100	cgc Arg	agc Ser	ctg Leu	att Ile	gac Asp 105	gat Asp	tcc Ser	gaa Glu	gag Glu	ggc Gly 110	ctt Leu	ggc	336
_	ej aaa		tag													348

WDN/SAS:gte 4/26/01 178-59010

P. 03

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May 1 2001

COMBINED DECLARATION AND POWER OF ATTORNEY FOR PATENT APPLICATION

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name.

sought	ventor (if plural na on the invention cr	mes are listed below) of the	intor (if only one name is listed b subject matter which is claimed SUS MACAQUE RHADINOVIR	and for which a patent is
	is attached hereto	s .		
	was filed on	as United States Applicat	tion No	
\boxtimes	was filed on Nov	ember 5, 1999 as Internatio	nal Application No. PCT/US99/2	26260.
	and was amended	d on (if applicable).		
	with amendment	s through (if applicat	ole).	
includi		nt I have reviewed and under mended by any amendment	rstand the contents of the above- referred to above.	identified specification,
in 35 U applies occurre continu	f Federal Regulation I.S.C. § 120 which It in the factor of the filing I for the filing of the filing I hereby claim for the filing I hereby claim for the filing I for patent of the filing I	ons, § 1.56. If this is a conti- discloses and claims subjec- lowledge the duty to disclosing date of the prior application. The priority benefits under inventor's certificate or of	ion which is material to patentabi inuation-in-part application filed it matter in addition to that disclo- e material information as defined on and the national or PCT inter- er Title 35, United States Code, § any PCT International application disclow and have also identified	under the conditions specified sed in the prior copending in 37 C.F.R. § 1.56 which national filing date of the 119(a)-(d) of any foreign on(s) designating at least one
applica	ition(s) for patent ountry other than the	or inventor's certificate or an	y PCT International application(filed by me on the same subject t	s) designating at least
	Prior Foreign A	Application(s)		Priority Claimed
	(Number)	(Country)	(Day/Month/Year Filed)	Yes No
applica	I hereby claim to ation(s) listed below	he benefit under Title 35, U	nited States Code, § 119(e) of an	y United States provisional
		60/107,507		er 6, 1998
	Ap	plication Number 60/109,409	~	Date r 20, 1998
	Ap	plication Number		Date
				Page 1 of

Fax:503-418-2719

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I hereby claim the benefit under Title 35, United States Code, § 120 of any United States application(s) or § 365(c) of any PCT International application(s) designating the United States, listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States or PCT international application in the manner provided by the first paragraph of Title 35. United States Code, § 112, I acknowledge the duty to disclose material information as defined in Title 37. Code of Federal Regulations, § 1.56(a) which occurred between the filing date of the prior application and the national or PCT International filing date of this application:

PCT/US99/26260	5 November 1999	Pending
(Application No.)	(Filing Date)	(Status: patented,
		pending, abandoned)

The undersigned hereby authorizes the U.S. attorney or agent named herein to accept and follow instructions from _____ as to any action to be taken in the Patent and Trademark Office regarding this application without direct communication between the U.S. attorney or agent and the undersigned. In the event of a change in the persons from whom instructions may be taken, the U.S. attorney or agent named herein will be so notified by the undersigned.

I hereby appoint the practitioners associated with the customer number provided below to prosecute this application, to file a corresponding international application, and to transact all business in the Patent and Trademark Office connected therewith:

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I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Scotion 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

Page 2 of 3

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WDN/SAS:810 4/26/01 178-59010

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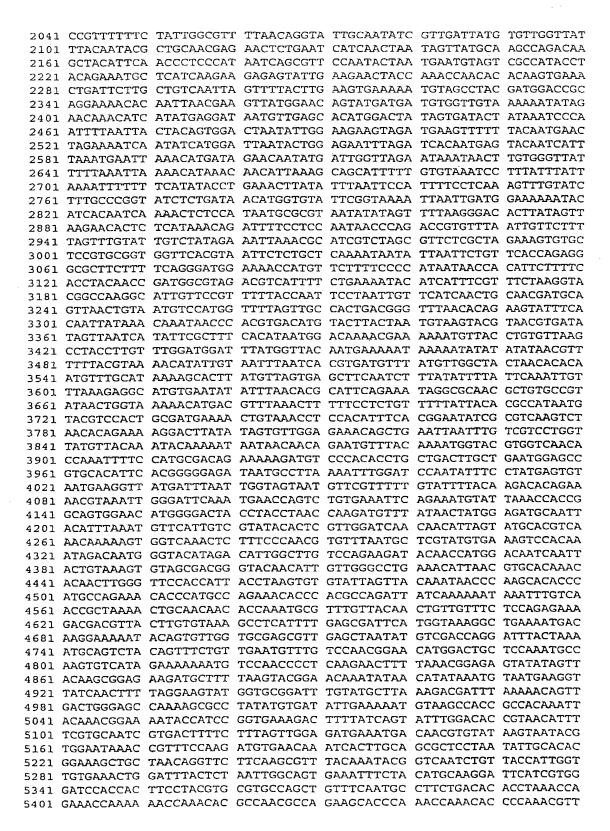
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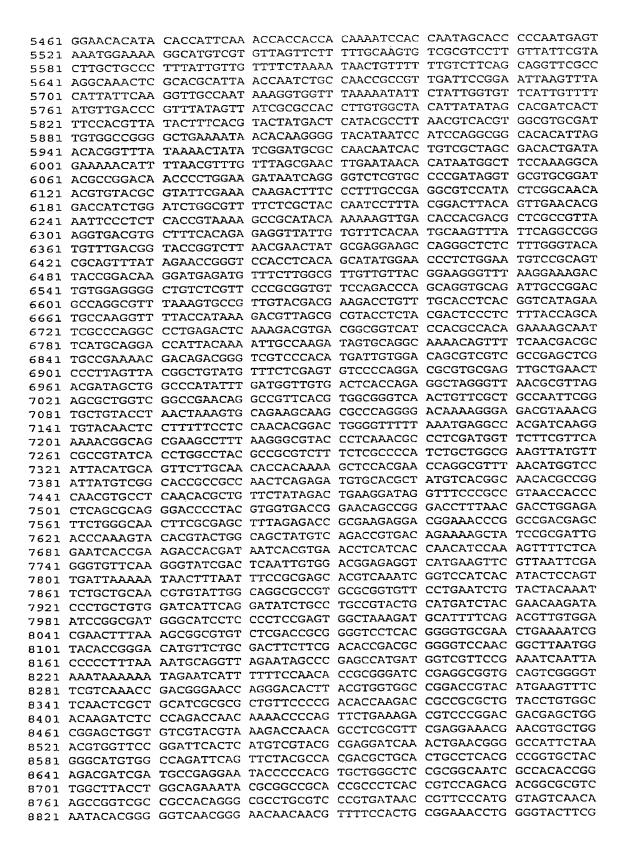
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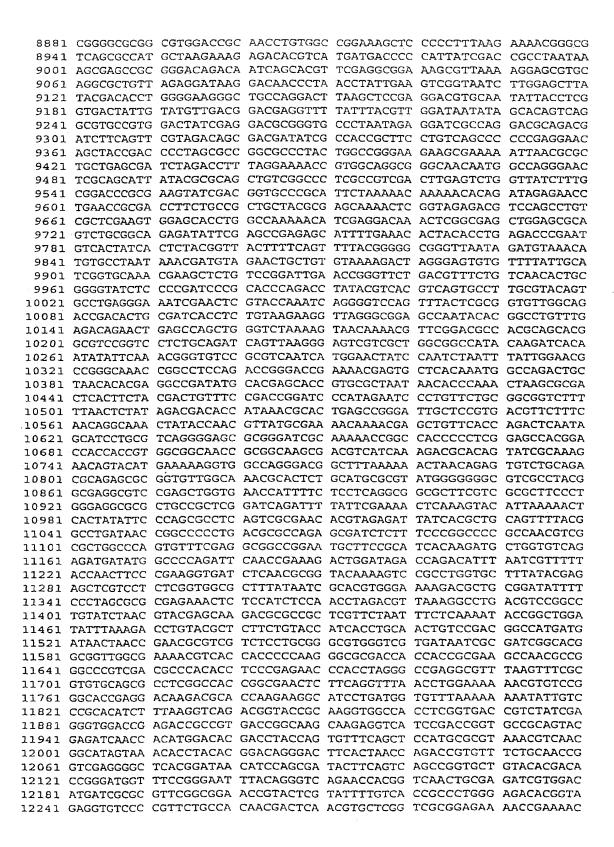
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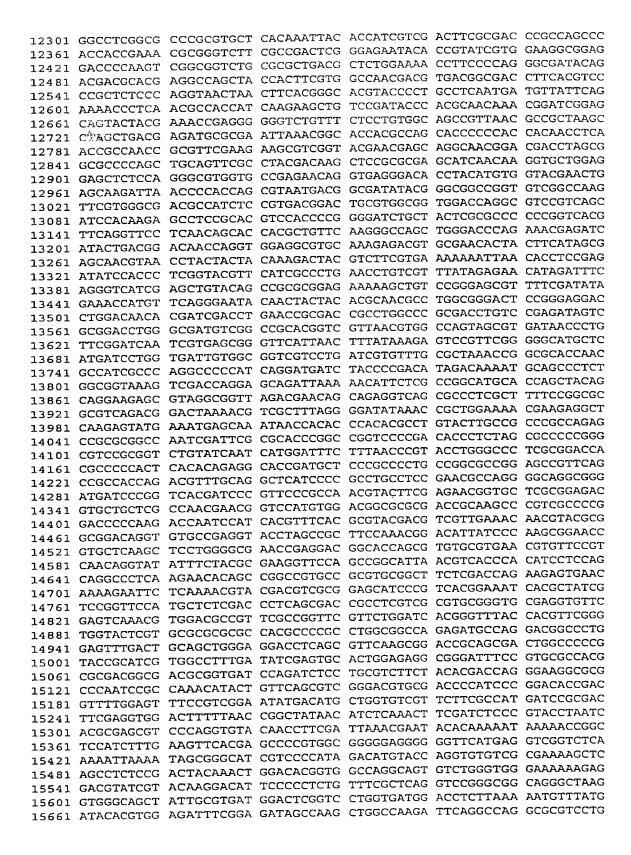
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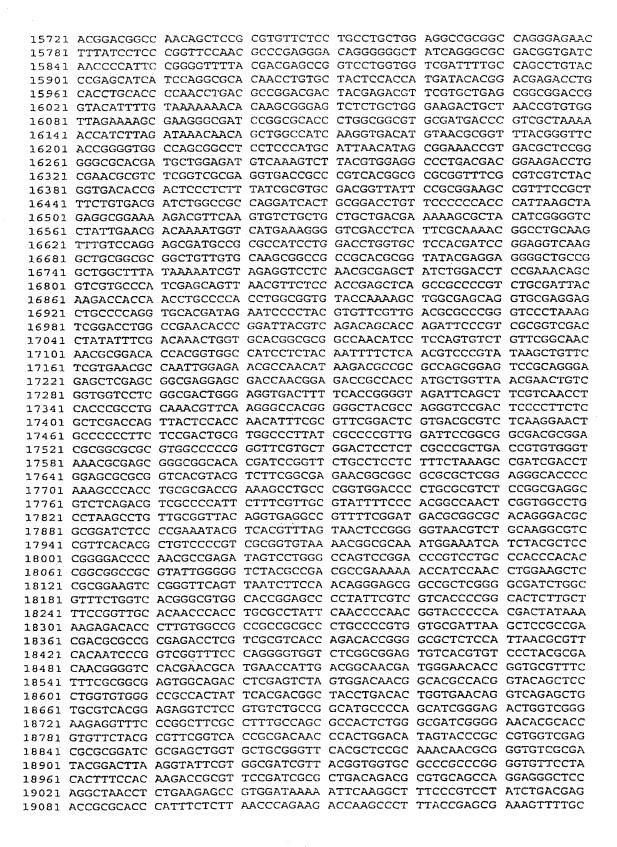
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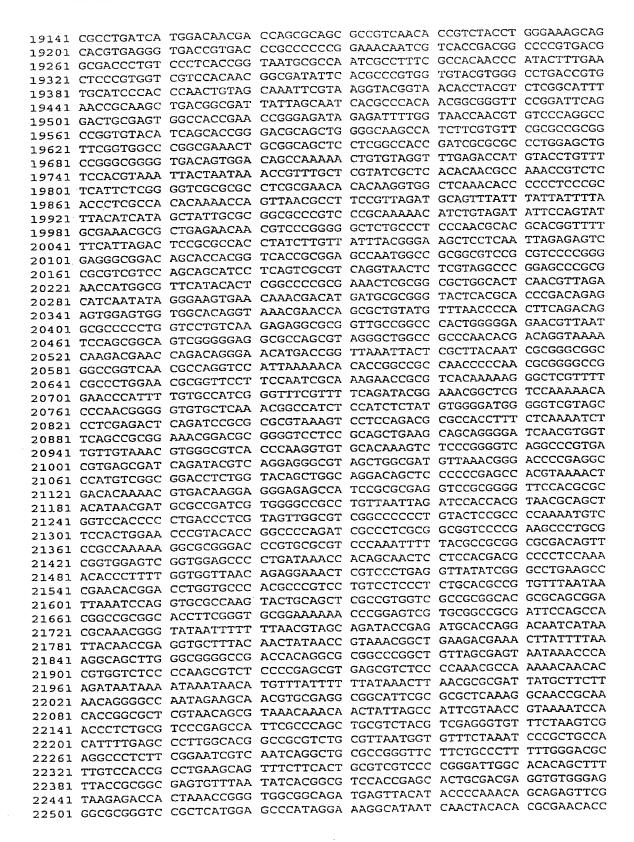
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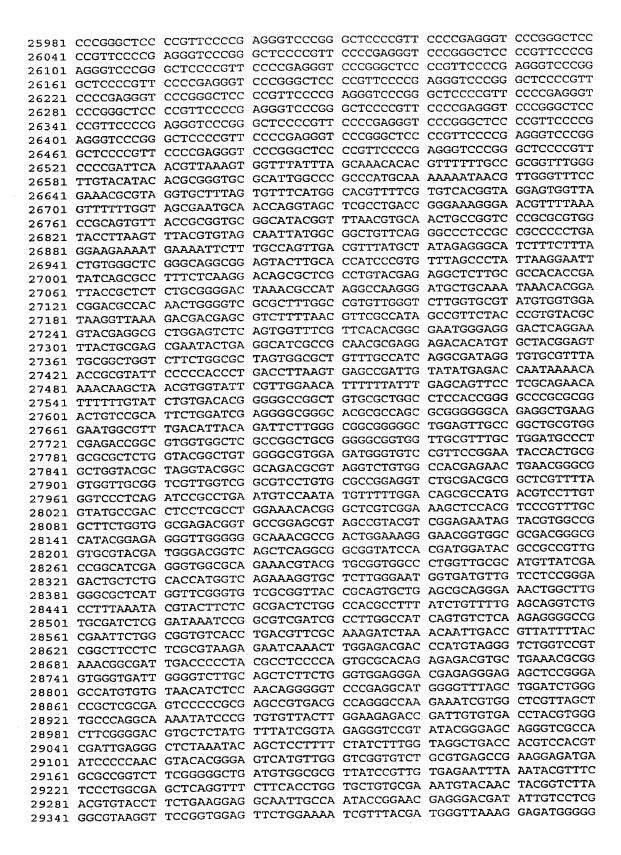
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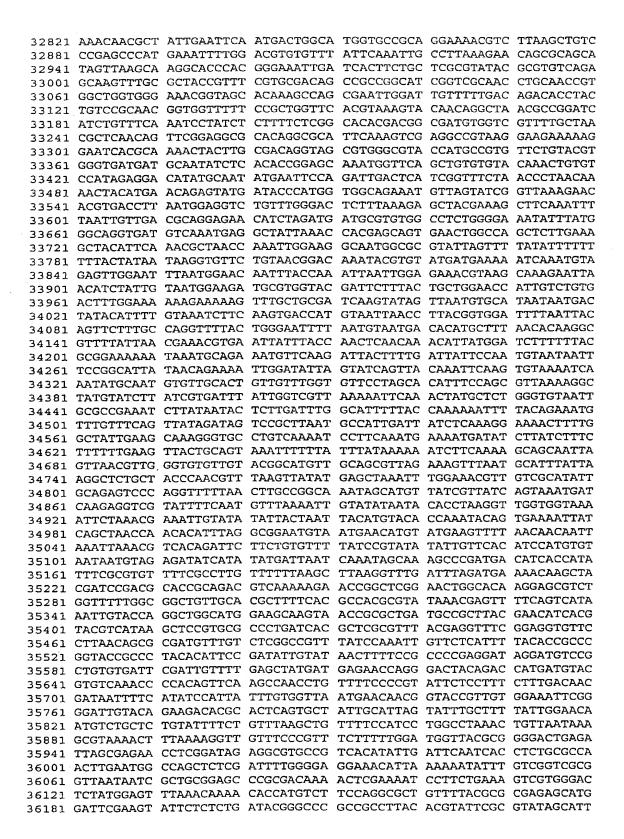


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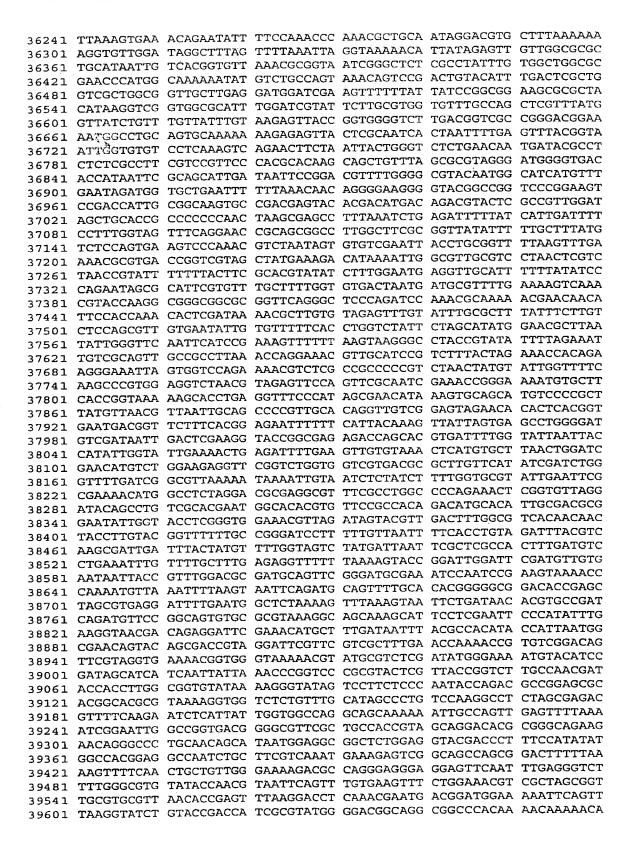
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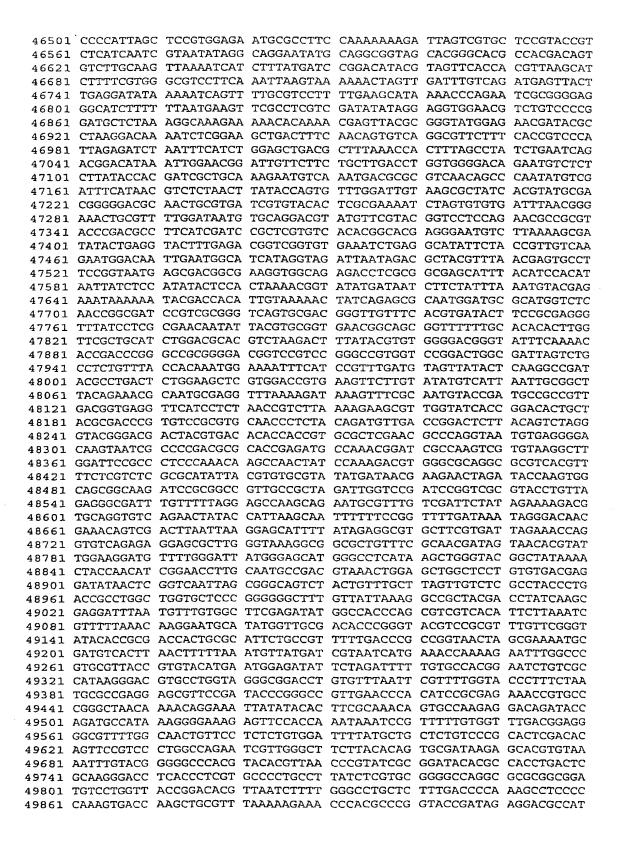


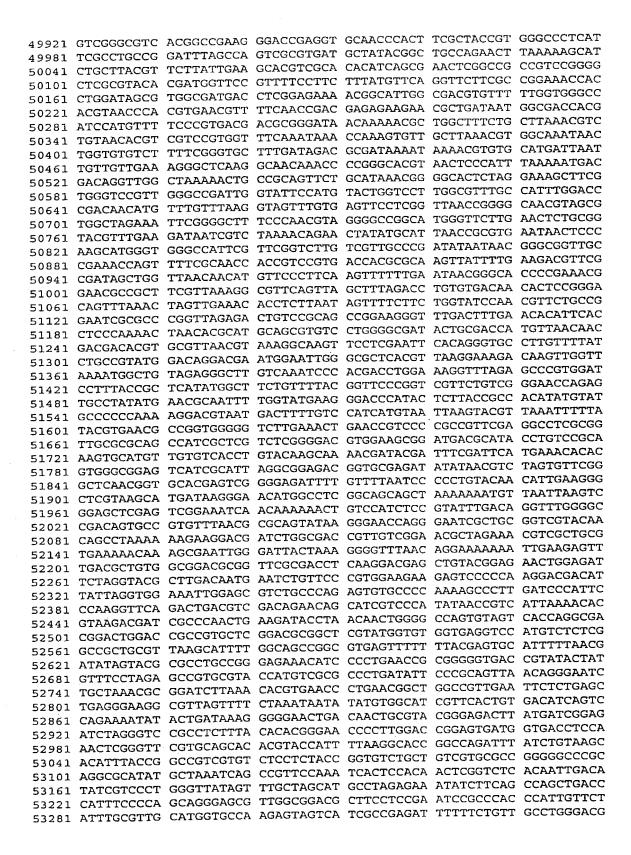
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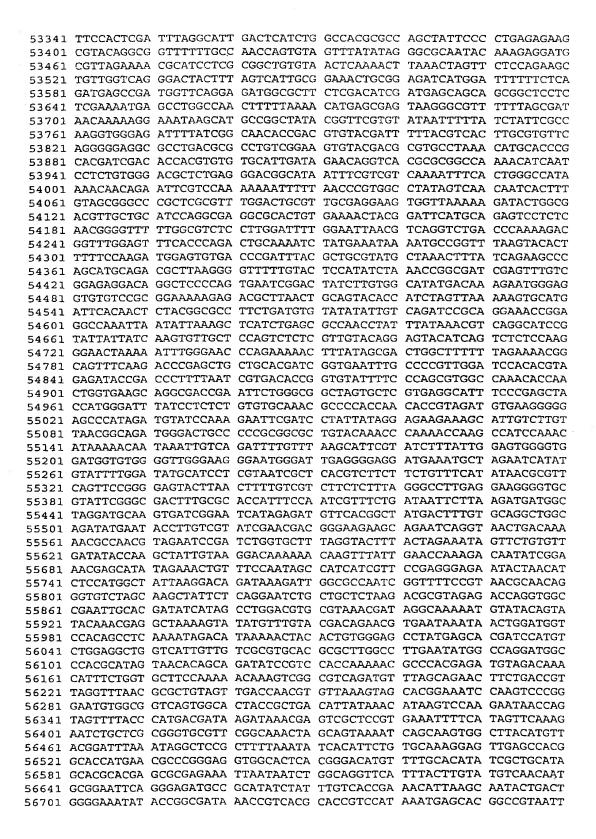
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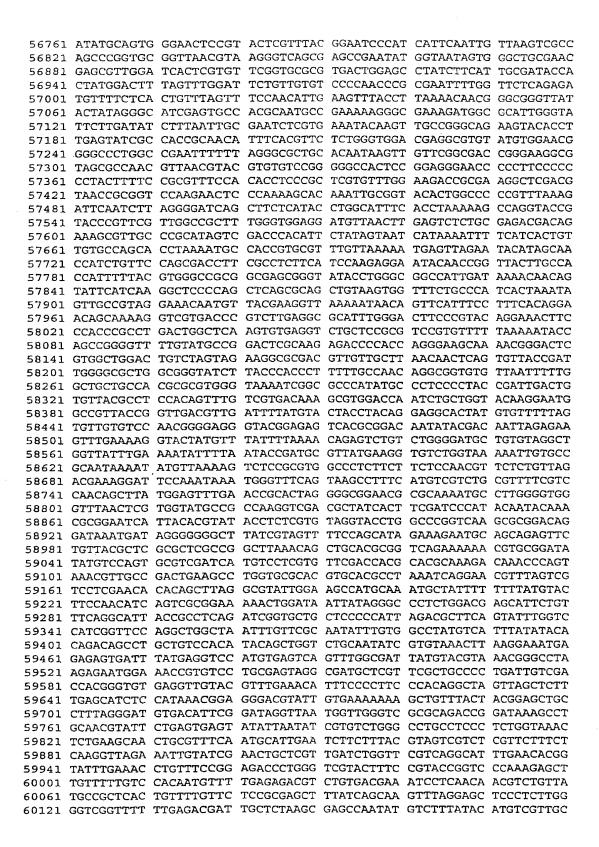
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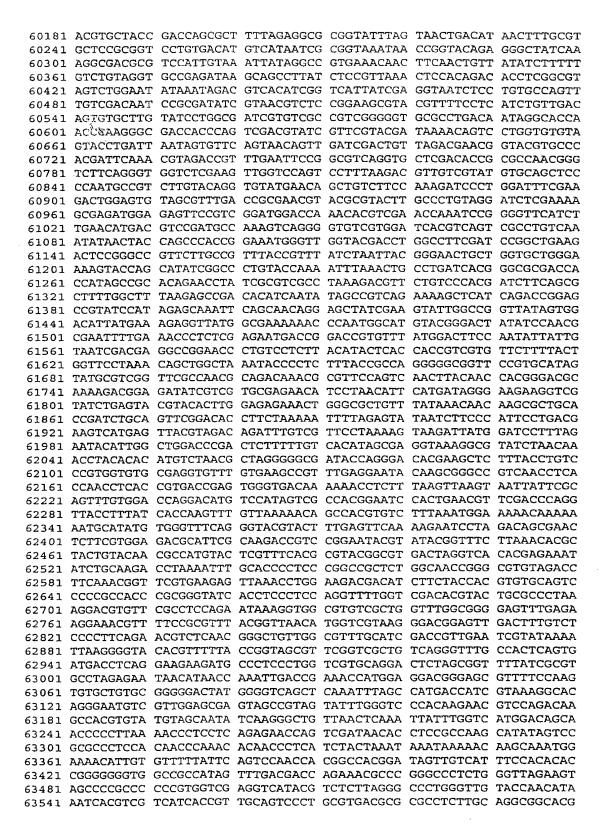
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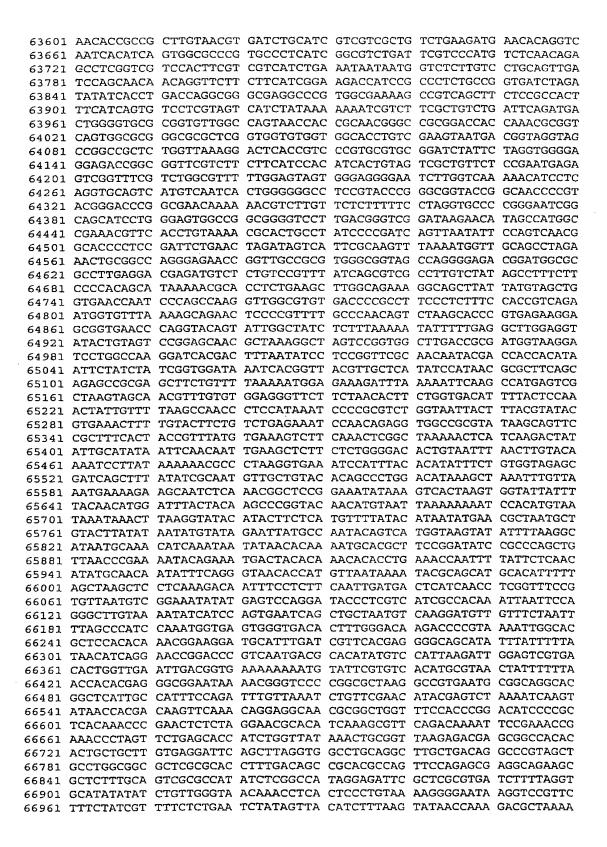








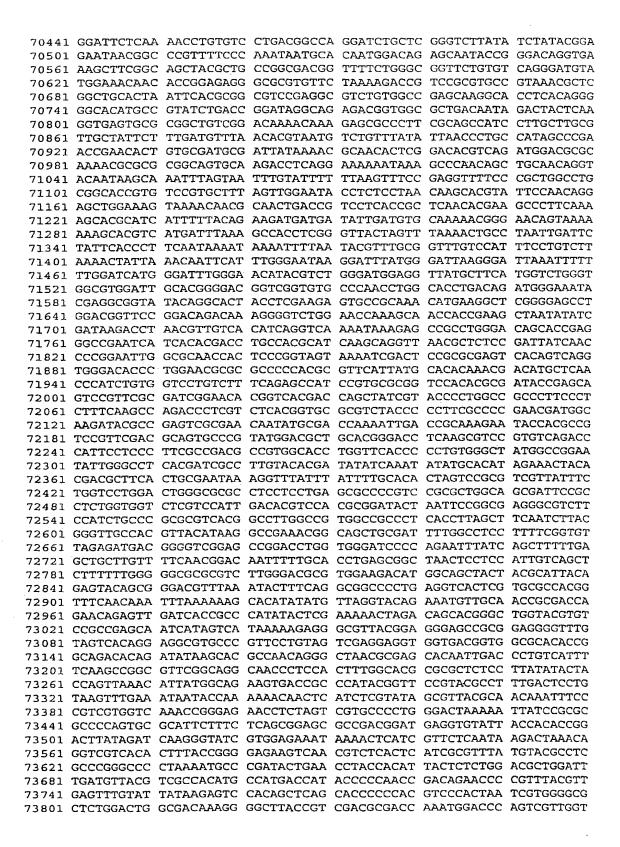


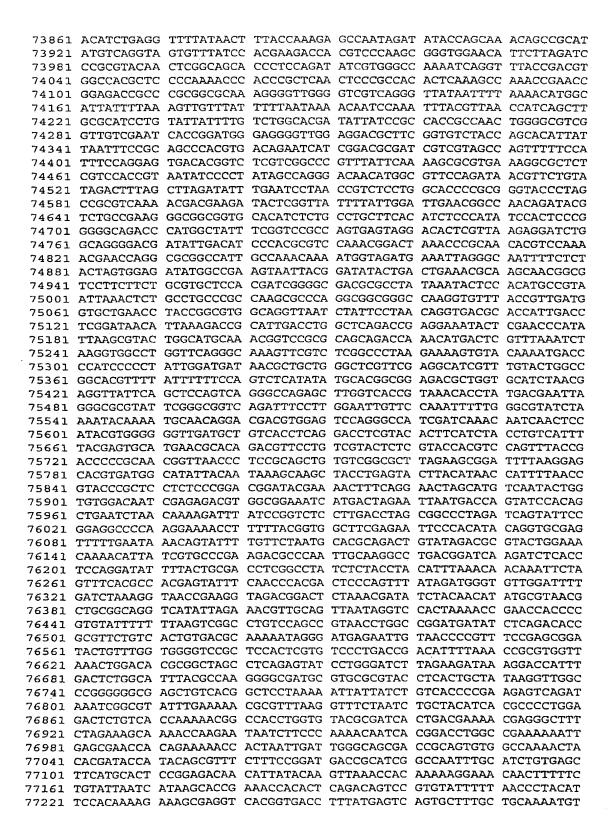




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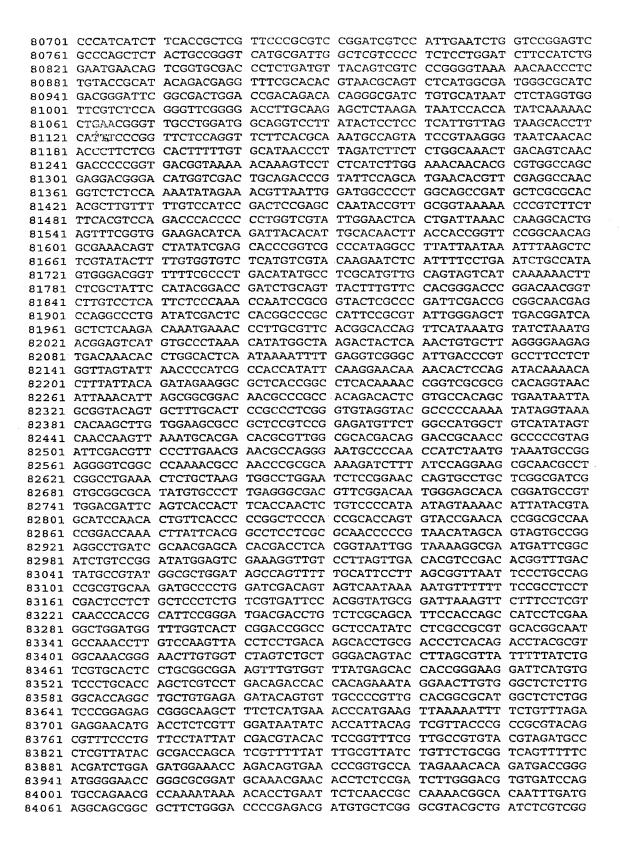


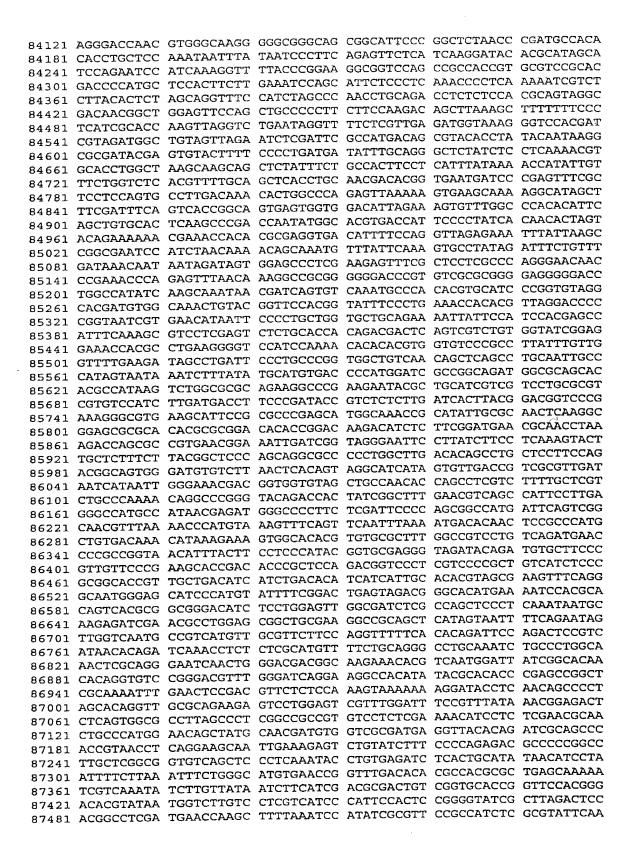


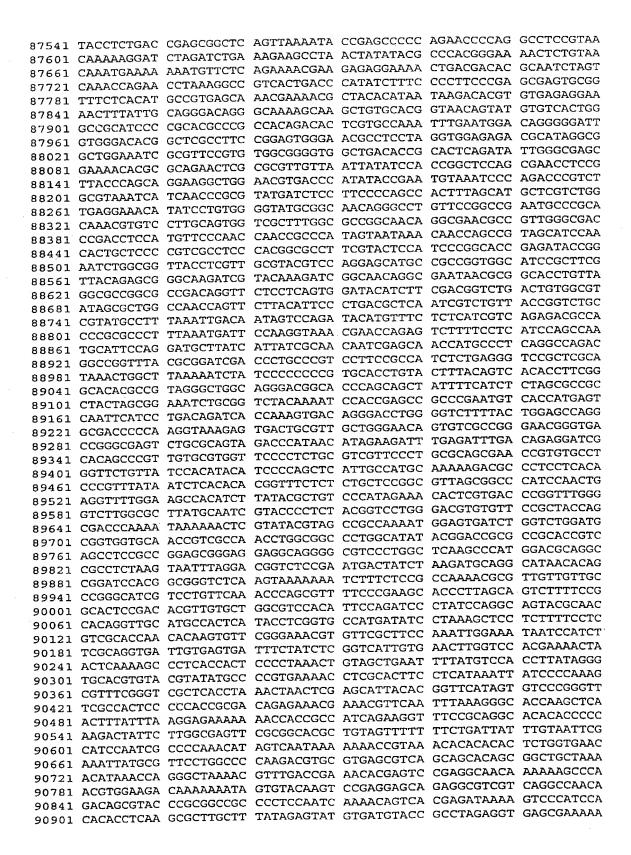
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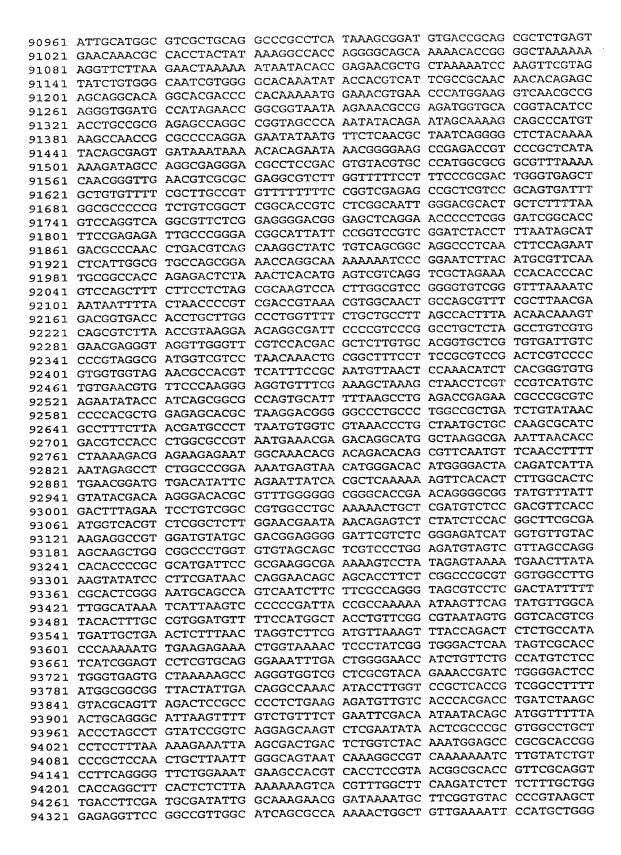
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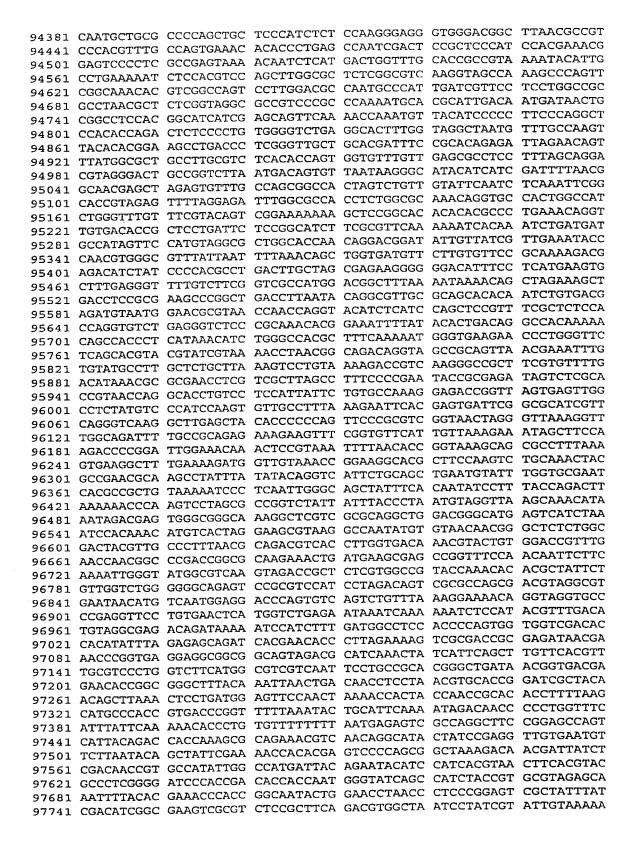




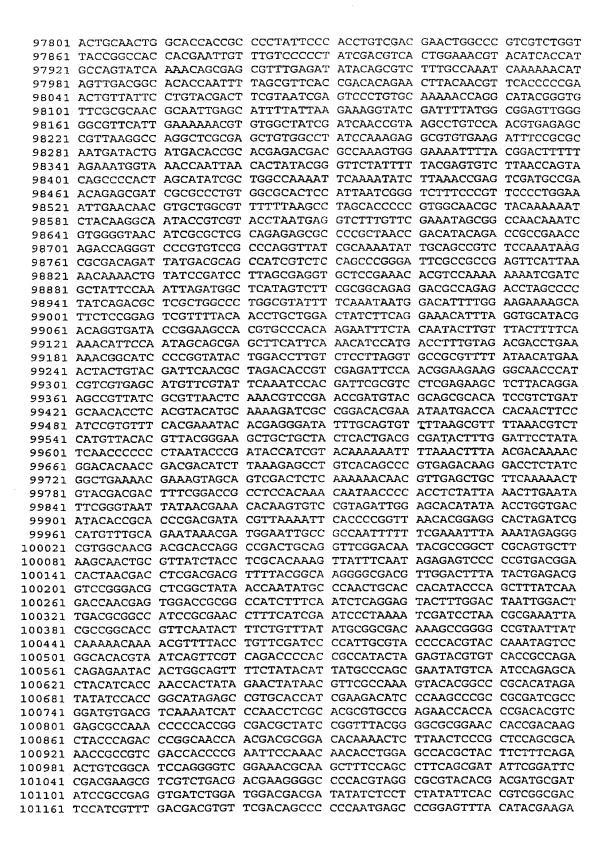


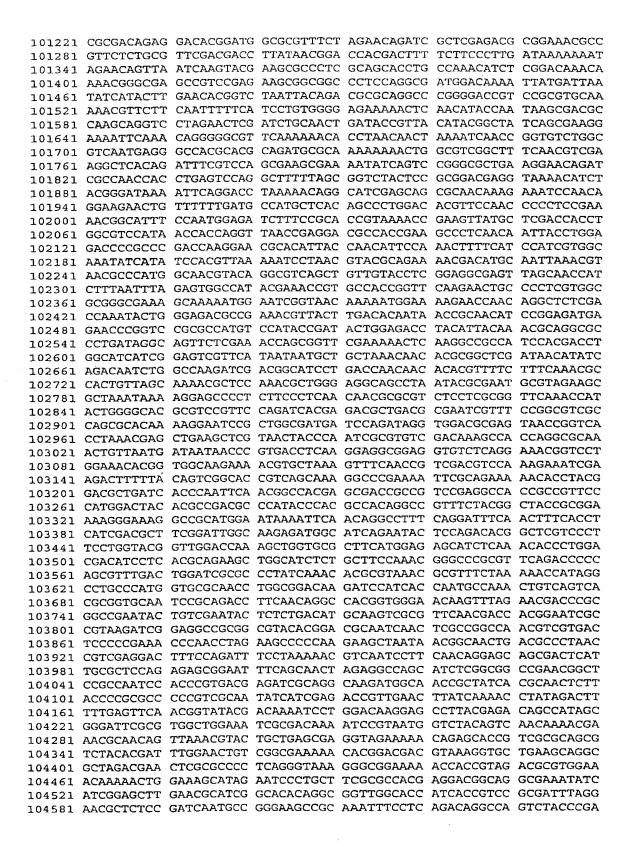










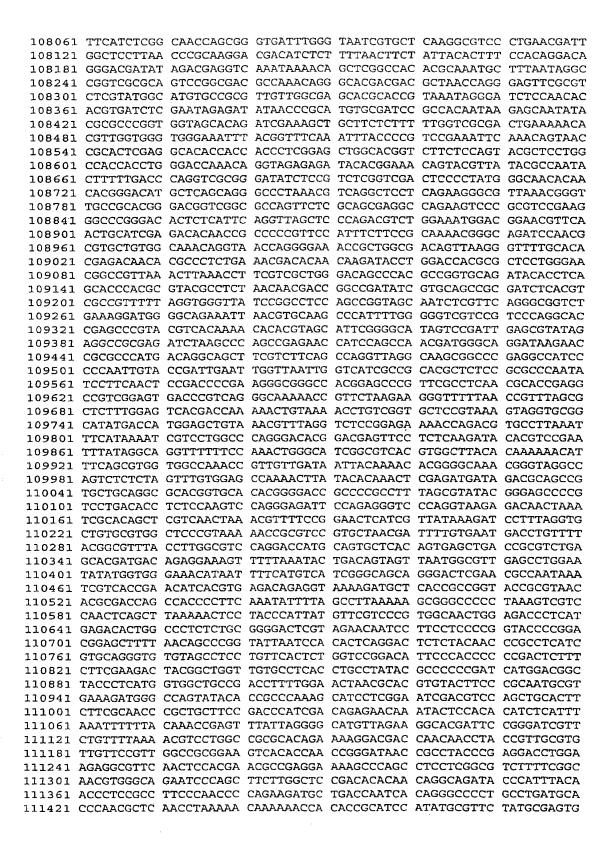




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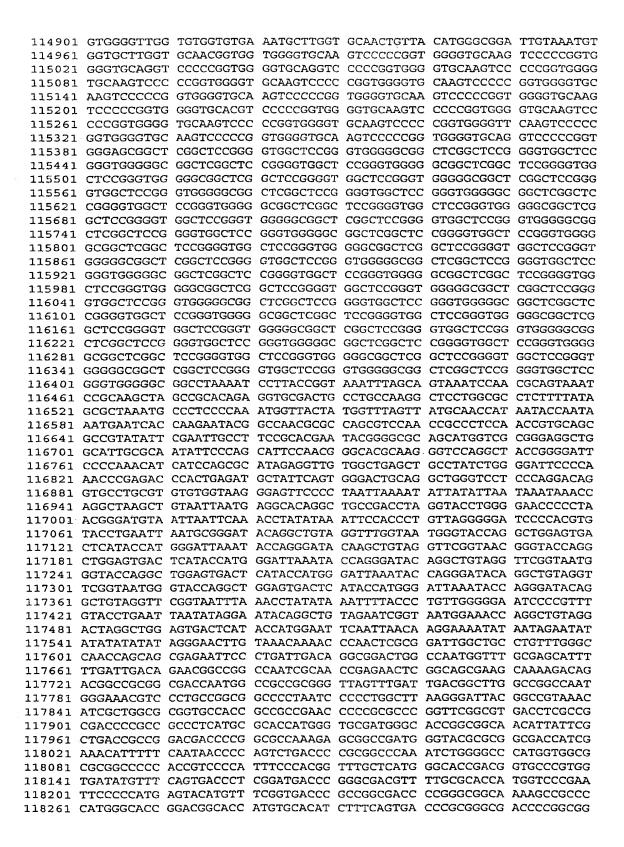




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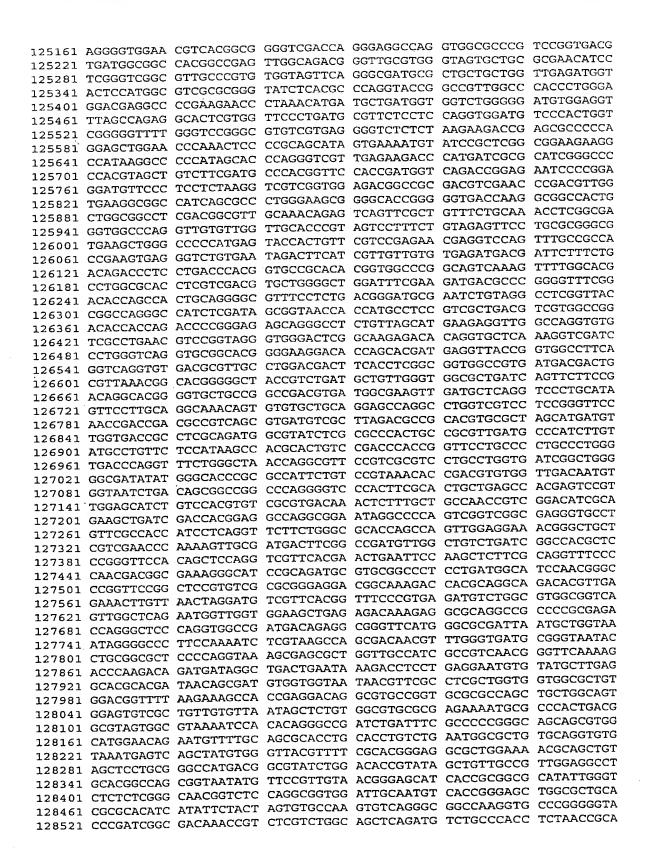
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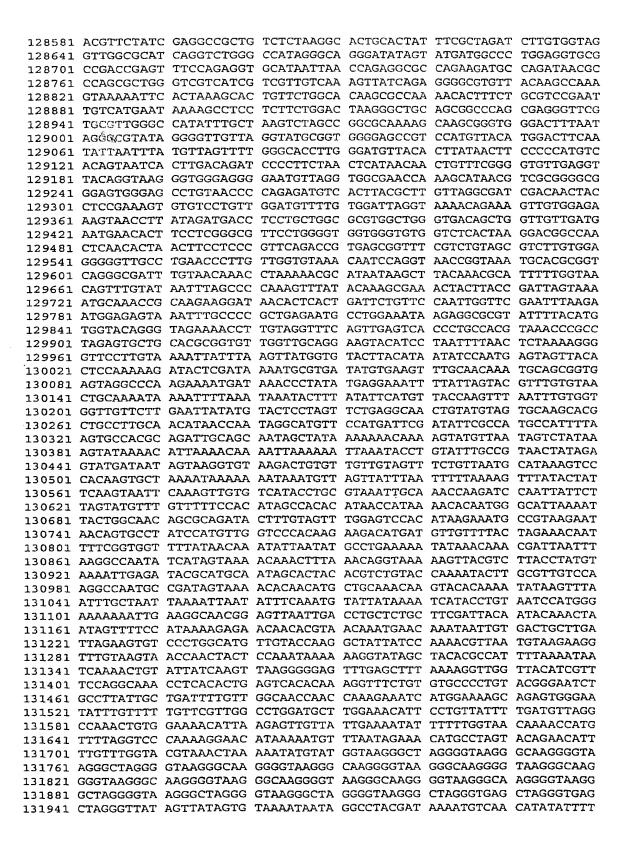
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SEO ID NO 2

CDS nucleotides 1353-2624

SEQ ID NO 3

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SEQ ID NO 4

CDS

complement (2692..3258)

/note="dihydrofolate reductase; ORF 2; similar to Kaposi's sarcoma-associated herpesvirus ORF 2"

SEQ ID NO 5

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SEQ ID NO 6 CDS

3676..5613

/note="complement binding protein; ORF 4; similar to
Kaposi's sarcoma-associated herpesvirus ORF 4"

SEQ ID NO 7

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SEQ ID NO 8 CDS

6045..9443

/note="ssDNA binding protein; ORF 6; similar to Kaposi's
sarcoma-associated herpesvirus ORF 6"

SEQ ID NO 9

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SEQ ID NO 10 CDS

9468..11528

/note="transport protein; ORF 7; similar to Kaposi's
sarcoma-associated herpesvirus ORF 7"

SEQ ID NO 11

/translation="MARELAALYAQLSALAVDLSLVIFADPRSIDGARILKTKTQIEN LNRDLLPLLREQNSVETSSLSLEVEHLAKNIEDKLGELERSLRQRYSSREHFETLHLR PECHYHSTVTFQFYGGGLIDVNMCLINDVELLCKRLGSVFYCIGANEALSGLNRVLTF LSTLRGISPIPHPDLYVTSVPCVQCLREIELVPNQGSSLLAVLADRHCDHLCKKVRAE PIHGLFETELSQLGLKVTKRSDATQHGVRSSADQLRESSLAAIQDHNIFKRVSASIME LSNLIYWNAGQTGLQTGTENECSQMARLLTHEADMHEHRALITPKLSATHFYDCFRPD PIESLFCGGLFNSIDDTINALSRDCSVTFFQQANYTNVMRKQNELFTRLNSILRQGSA GSQKPATPSEPRTTTVAATAASDVIKDAQYRKEQYMKKVARDGFKKLTECLQTQSAVL ANALCMRVWGGVAYGEASELVNHFLLRRRFVALPWEARCRSDQILFENSKYIKNSLYS QRLSREHVEIITLQFYGLITGPLTRQSDLFPGPANVALAQCFEAAGMLPHHKMLVSEM IWPQIQPKDWIDQTFNRFYQLPEGDLNAVQKSAWCFIRELVLSVALYNRTWEKTLRIF SLAREKLSISNLDVKGLTSGLYLTYEQDAPLVLISQNTGWIFKDLYALLYHHLQLSDG HDDN"

SEQ ID NO 12 CDS

11515..14004

/note="glycoprotein B; ORF 8; similar to Kaposi's
sarcoma-associated herpesvirus ORF 8"

SEQ ID NO 13

/translation="MMITNRTRRLLRAWVVIIAIGTAVGENVTTPKGATTTAKPTPGP STPTPPENPPRAEAFKFRVCSASATGELFRFNLEKTCPGTEDKTHQEGILMVFKKNIV

PCT/US99/26260

PHIFKVRRYRKVATSVTVYRGWTETAVTGKQEVIRPVPQYEINHMDTTYQCFSSMRVN VNGIVNTYTDRDFTNQTVFLQPVEGLTDNIQRYFSQPVLYTTPGWFPGIYRVRTTVNC EIVDMIARSAEPYSYFVTALGDTVEVSPFCHNDSTCSVAEKTENGLGARVLTNYTIVD FATRQPTTETRVFADSGEYTVSWKAEDPKSAVCALTLWKTFPRAIQTTHEASYHFVAN DVTATFTSPLSQVTNFTGTYPCLNDVIQKTLNATIKKLSDTHATNGSEQYYETEGGLF LLWQPLTPLSLADEMRELNGTTPAPPTTTSTANRVRRSVGTNEQATDDLAAPQLQFAY DKLRASINKVLEELSRAWCREQVRDTYMWYELSKINPTSVMTAIYGRPVSAKFVGDAI SVTDCVAVDQASVSIHKSLRTSTPGICYSRPPVTFRFLNSTTLFKGQLGPRNEIILTD NQVEACKETCEHYFIASNVTYYYKDYVFVKKINTSEISTLGTFIALNLSFIENIDFRV IELYSRAEKKLSGSVFDIETMFREYNYYTQRLAGLREDLDNTIDLNRDRLARDLSEIV ADLGDVGRTVVNVASSVITLFGSIVSGFINFIKSPFGGMLMILVIVAVVLIVFALNRR TNAIAQAPIRMIYPDIDKMQPSGGKVDQEQIKNILAGMHQLQQEERRRLDEQQRSAPS LFRRASDGLKRFRGYKPLENEEAQEYEMSK"

SEQ ID NO 14 CDS

14122..17166

/note="DNA polymerase; ORF 9; similar to Kaposi's
sarcoma-associated herpesvirus ORF 9"

SEQ ID NO 15

/translation≈"MDFFNPYLGPRGPRPHSHRGTDAPAPAGAGAVQPPPDVCRLIPA CLRTPGAGGMIPVTIPFPPTYFENGARGDVLLANERSMWTARDRKPVAPDPQDQSITF HAYDVVETTYAADRCAEVPSRFOTDIIPSGTVLKLLGRTEDGTSVCVNVFRQQVYFYA KVPAGINVTHILQQALKNTAGRAACGFSTRRVNKRILKTYDVAEHPVTEITLSSGSML STLSDRLVACGCEVFESNVDAVRRFVLDHGFTTFGWYSCARATPRLAARDARTALEFD CSWEDLSVOADRSDWPPYRIVAFDIECTGEAGFPCATRDGDAVIQISCVFYTTREGAP NPPNILFSVGTCDPIPDTDVLEFPSEYDMLVSFFAMIRDFEVDFLTGYNISNFDLPYL ITRASQVYNLRLNEYTKIKTGSIFEVHEPRGGGGGFMRSVSKIKIAGIVPIDMYQVCR EKLSLSDYKLDTVARQCLGGKKEDVSYKDIPPLFRSGPGGRAKVGSYCVMDSVLVMDL LKMFMIHVEISEIAKLAKIQARRVLTDGQQLRVFSCLLEAAARENFILPVPTPEGQGG YOGATVINPIPGFYDEPVLVVDFASLYPSIIQAHNLCYSTMIHGRDLHLHPNLTPDDY ETFVLSGGPVHFVKKHKRESLLGRLLTVWLEKRRAIRRTLAACDDPSLKTILDKQQLA IKVTCNAVYGFTGVASGLLPCINIAETVTLRGRTMLEMSKSYVEALTTEDLRTRLGRE VTARHGARFRVVYGDTDSLF1ACDGYSAEAVSAFCDDLAAR1TADLFPPP1KLEAEKT FKCLLLLTKKRYIGVLLNDKMVMKGVDLIRKTACKFVQERCRAILDLVLHDPEVKAAA RLLCKRPPHAVYEEGLPAGFIKIVEVLNASYLDLRNSVVPIEQLTFSTELSRPVCDYK TTNLPHLAVYOKLASRCEELPQVHDRIPYVFVDAPGSLKSDLAEHPDYVRQHQIPVAV DLYFDKLVHGAANILOCLFGNNADTTVAILYNFLNVPYKLFS"

SEO ID NO 16

17261..18511

/note="unknown; ORF 10; similar to Kaposi's

sarcoma-associated herpesvirus ORF 10"

SEQ ID NO 17

translation="MLVNELSVVLGDWEVTFHRGRFSFVNLTRLQTFKGHGGYARVRL PFSLDQLLHQHFAFGLVTRLKELPPFSDCVALIAPLDSGGDADAARVAPGFVLDSSRP LTVWVNASGRHTIRFCLLFLKPIDLERAVTYVFGENGGARSEGTPKPTCATESLPGGP LRVSGEASQTSPHSFVAYFPTANSVACLSLLRLQVRPFSDDAAHRDARISPKYVTFSN SGGNVCKASVHTLSPSRCKTAQMEIIYAPGDPNAEIVLGQSGPVLPTHTGGRVLGVYA DAEKTIQPGSSAEVRVQLIFQQGAAARGDLAFLVTGVAPEPLFVVTPALLLSGCTTHL RLFNPNGTPTTIKRDTLVAAAAPCPVVRLSSADDAPRDLVASPDTGALSINAFTIPVG FPGVVSAECHVSLRDNGVHERMNH"

SEQ ID NO 18

CDS

18520..19749

/note="unknown; ORF 11; similar to Kaposi's

sarcoma-associated herpesvirus ORF 11"

SEQ ID NO 19

/translation="MGTPVRFFRGEWQTSSLVDNGTPRYSSLVWAATIHDGYLTLVNR SELCVTERSPCLPACPSIGRLVGKRFPGFAFASATLGDRGTRTVFYAFGHRDNPLDIV PAVVERADRELVLRVHAPQTTRVSRYGLKVFVAIVTVVRPPGVFLHFPQDRVPIALTD ACSQEGSRLTSEEPWIKIQGFPVLSDETAHPFLLTQKTKPFTERKFCRLIMDNDQRSA VNTVYLGKQHVRVTVTRPPETIVTDGPVTATLSLTGNAPIAFRHNPYFELPWSSTTAI FTPVVYVGLTVCIPPNCSKFVRYGNTYVSAFNRKLTAIISNHAHNGGFRIQDCEWPPN

 ${\tt REIEILVTNVSQAPVYISTGTQLGQAIFVFAPRFGGPAKLRQLLGHRSRALELPGGVTVDSQKLCRFETMYLFST"}$

SEQ	ID	ИО	20	CDS	complement (1992120544)
SEQ	ID	NO	21	,	/note="R2; similar to IL-6" /translation="MFPVWFVLFYLSCWAASPTLAPPPTAAGINVLPQWAGNRASLDR TRGRLSEVGLNIQRWFVYLCHHSTLCRVREYPRIMSFVHFPILMSNVECQRREFRGAE CMNAMVRGLRAYESYLTRLRMLLDDAPGDADAAAIGSAVTVVLSALDSLIEELPVNNK IGGAESNEKTVRALGGQSPRDVVLSAFRILEYLQMFLRDGRRAIAMM"
SEQ	ID	ИО	22	CDS	<pre>complement(2077721778) /note="thymidylate synthase; ORF 70; similar to Kaposi's</pre>
SEQ	ID	йО	23		sarcoma-associated herpesvirus ORF 70" /translation="MIVLVHLGICYVKKIIPVCVAGIAAARLRVFSAPEGAAAVRCAC RGDHGELQYLAHLDLIIKHGVQREDRTGVGTRSVFGLQARYNLRDEFPLLTTKRVFWR GVVEELLWFIRGSTDSTELSRRGVKIWDAHGSRAFLAAQGFGDRREGDLGPVYGFQWR HFGAEYRGADANYEGQGVDQLRYVVDLINRRPHDRRIVMCAWNPADLARMALPPCHVL CQFYVARGELSCQLYQRSADMGLGVPFNIASYALLTYLIAHVTGLTPGDFVHTLGDAH VYNNHVDPLLLQLRRTPRPFPRLKILRKVARLEDFTRADLSLEGYDPHPHIEMEMAV"
SEQ	מד	MO	24	CDS	complement(2224522592)
DDQ		210	2.		/note="R3; similar to Kaposi's sarcoma-associated
SEQ	ID	ио	25	/t	herpesvirus K4 viral MIP" ranslation="MRGLFVCVFFAVFACVVDYAFPMGSMSGPAPELCCLGYVTHLPP PGLVVSYSHTSSOCSVDAVILNTRRGKKLCANPGDDAVKKLLQAVDKRPKKGRRTRRS
					LIDDSEEGLGSGI"
SEQ	ID	NO	26	CDS	2684627409 /note="Bcl-2 homolog; ORF 16; similar to Kaposi's
SEQ	ID	ио	27		sarcoma-associated herpesvirus ORF 16" translation="MAAVQGPPPPPEEENENSLPVDVYAIEGIFLYCGLGQAEYLHHP VFSPIKEFISAFLKDSARLYERLLRHTDYRSLRGLNAIGQGMLQINTDGRHNWGRALA VLGLGAYVVDKVKDDERLLTFAIAVLPVYAYEALESQWFRSHGEWEGLRNYCERILRH RRNARRHMCYGVAAGLLALVALFAIRR"
SEQ	ID	мо	28	CDS	complement (2751529125)
					<pre>/note="capsid protein; ORF 17; similar to Kaposi's sarcoma-associated herpesvirus ORF 17"</pre>
SEQ					translation="MTPVYVGGYVDVVSLPKIEKELYLEPSIVATLLPYTDPLPINIE HVPEAHVGHTIGLFQVTHGIFCLGKLTSHDFLALASRLAGDSRAAQIQLNPMPRDPLL EMLHTWLPELSLSSLHPEELQDPNHPPAFQHVSLCALGRRRGSIAVYGPDPTWVVSKF DSLTREEAGKITVNCLDLCERQVTPPEFAAPLETLMAKAIDAGFIRDRTDLLKTDKGV ARVARSTYLKASQFPCAQHCGNRDTRTMSALPEDNITIPKSTFLTMVQSSLDNMRNQG HRTYVSAPPSMPATAAYPSWIPPPELTVPSYAPPVAPPFPFQSAFAPQPSPYAATYYS PTYGYAPAPSRHQKRKRDVELSDEPVFPGEEVGIHKDVMALSKNILDIQADLRDLKRA ASQTSGAQDADQRPQPPPVQFSWPQTYASAPYLAYQPQWYSGTDTHLHAPQPYQSAQG IQQTQPPPPQPASHHAGLATQPATPAPAAQESVMSNAIPSASAPRAGACPPLDPECGQ SARAPVEASAQPAPVSQIQKMFCEELLK"
SEQ	ID	ŇO	30		2899829897 /note="unknown; ORF 18; similar to Kaposi's
SEQ	ID	NO	31	; ; I	sarcoma-associated herpesvirus ORF 18" /translation="MFIGRGSVYGSRVATIEGSKYSSFSIFGRLTTSTYPPTYTGVML GRCLREPKEMSAGLRGLMWRVIRCENLNTFLPGELRFLHLVLCEMYNYGLNVYLLKEA IANTGTRDDIVLGRKVPVEFWKIVYDGLKEMGVSDATLLSETKRGALWLYFNGRPCLL KGLGDYVFCQLGLSHSVRVVPENLTDGNYLYNLGSVIPCRLLVALSYCLAFWGHADHE PWVRLFAGKIFILYLIISGHIMPRKSILEQVGTSGYGGFVEAVCRDVRAVHGIPAWDF

ASAAPALTSQQTDYLFAFNNSVV"

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SEQ ID NO 32 CDS

complement (29905..31548)

/note="tegument protein; ORF 19; similar to Kaposi's

sarcoma-associated herpesvirus ORF 19"

SEQ ID NO 33

/translation="MRTSEKCCMRYPRKPARQITATFWAPHPNNVLFIHKPSLIEERR NAFVMRNQQLALRVHTLRKNLLRLELDNVLQTHQRETEVVMRDLETIQNMVGDLRSPG RETANAQTSLNPQPKIAPQTHGDAFVVTIAPGDPGFTVNQDLRLELLPSLYMNQNQWL PQYGPWYSSLTDNAMQRRVFPRDLRGTTNFQNSTSLKLMSAVISTAASITQDFYADVR NVSDTQAALCLLNGYYCHRTGTPLPPTRNGLWDNLGTKLATLVSHLKQNTKGLGFEFT YSNPRQRASLAPLNKETKYNADFFTNHVIYATLAQSGLLPGSKNPGTGQPPGPDLVYI LATTLFSEDVPPFQAYQWNLRAGLSALGCLVLVYVLLELAQITPRSPHRRLNLASLLG GRFSKVEDPSGSKQYLKKGQLFDFLTENYISPILSRAPDAPTSFLFPGAYLAALEAKA ISHLKHTRPFVNLTGSRFNEIFDILNQKLTFRDAGSLIQAQTSLRLTAEEGLAAILSH PSPPGLAHEIMKSQFGVYDDYDRVYFLVLGYLPVATSVV"

SEQ ID NO 34 CDS

complement (31043..32095)

/note="unknown; ORF 20; similar to Kaposi's

sarcoma-associated herpesvirus ORF 20"

SEQ ID NO 35

translation="MAFANQCKHVATLEALPASRKRAGTRAHLAVYRRLIKHRSLDDI LKFLSIRPTLRATKNVKFRIFFEVSLGRRIADCVLTVNSEHQKTCYVIELKTCLSAAV FPGNAIKISQRWQGLHQLTDSVAYIGRAAPRGHENWSVRPWLLFKNQKTLKTIHTESS AFPPTFINTTSAALNGFFSQWEDAHVRKMLYEIPTKTSAANYRNFLGPPSKQRSVYSQ TISDRRKKKRVCDAKSTAGAKGSHAAKKPAPARTRQRAANAPTGNRSGHARPRNNSKH GRGSAVPGQGNRQCPNITKPATQNRPADTWRRVRCHNSPRRPGIHGKPGSPSGAPAKP VHEPKPMAATIRAVVO"

SEQ ID NO 36 CDS

32094..33767

/note="thymidine kinase; ORF 21; similar to Kaposi's

sarcoma-associated herpesvirus ORF 21"

SEQ ID NO 37

/translation="MAEGSGFGDELVRQMRDRKPRWDESSDDTDDVDTESTDLEYDD VFPVVDTHGLMSPGSQNYDVPTSPSGTPWELLHPDALYAHPRCPPKRAVVPGGGARPK VSAFSARLQYVGRQSFGDRETRQLTGAQFSSESEHEYAEIPERTTTRPVESGDKRNFT SGRRGAISGPSSTKPSHGAGLTRKTKTSLSVSLKNLLRIKDDDAKVDVPRPVTVPVHL MQPHPMTEYRNAFLIYLEGVMGVGKTTLLNSMTGMVPQENVLSCPEPMKFWTCVYSNC LKEQRSIVKQGTHGKLITSARVYACQSKFALPFRATAAGIGRNLQPWLVGNGSTKPAN WIVFDRHLLSATVVFPLVHVKYNRLTPDHLFQILSLFSAHDGDVVVLLTLNSSEAHRR IQSRGRKEEKGITQNYLRQVAWAYHAVFCTWVMMQYLTPEQMVQLCVQTVSIEDICNM NSRLTHRFLTLTKLHEQSMIPMVAEMLVSVKEHVTLMEVCLGLFKELRKLQILIVDAG EHLDDACGLWGNIYGQVMSNEAIKPRAVNWPALESYIQTLTKLEGNGAY"

SEQ ID NO 38 CDS

33754..35868

/note="glycoprotein H; ORF 22; similar to Kaposi's
sarcoma-associated herpesvirus ORF 22"

SEQ ID NO 39

/translation="MARISFIFFTIIRCSVTDKYVYDEKSNVELEFNGTIYQINWRN VSKELTSIVMEDAWYDSLLLEPLSVTLEKRKSLLRSSIVNVHNNDYTFCKSSSDHVIN LTVDFNYSSLPGFTGNFNVMTHALTQGVLLTKRELFTNSTNIMDLFYAEKINAEMFKI TFDYSNVIISGIITENWILVSVTNSSVKSNMQCVALLFGVPSTFPALKGYVSYRDLLV VKNSNYALGVIAPKSYNTLDLAFLPKNFTEMFVSVIDSPLNAIDYLKGKLLAIEAKGA CQNPSNENDILSFFFEVTAVNFLFIKNLQKQQLVNVGCVVRHVAALESLMHLLRLCYP TFKLYELNLETLSHIAESQVFNLPANSMLSLSVNDQEVVFSMFKIVYNTPKVGGKILN EIVYITNYMYTKYSENYQLTNTFRRNVMNMYEVLTTIKLNVTDSSVFYPYILFTSMCN NVEISYMINQIAKPDDITIFRVFSPCFLSLRFDLDENKLRSDAPQTSKRTGSELAQGA SGFWRLLHAFHATRINEFSVINCTRLAWKQVTALMPLTNITYVISSVRPDHARVYEVS EVFLNSAMFVSAVYPNCSHFTPPGTALHIPILYNFSAPRIGCPLCDSIVLSYDENQGL QTMMYVSNPTVQANLFSPYSPFFDNDNFHIHYLWLMNNGTVVEIRGLYRRHALSAIAL VFAFIGTMSALYFLFKLFSILA"

PCT/US99/26260

SEQ ID NO 40

SEQ ID NO 41

CDS

complement (35865..37073)

/note="unknown; ORF 23; similar to Kaposi's

sarcoma-associated herpesvirus ORF 23"

/translation="MIKISDLKARLVGGAVQLSNGEYVCHVVYSSALAAMVGLPGPAV PLPLLFKKFSTIYSNMMPLYAPKRPELSMLRIMVSPHPYALNSCLCVGTDEGERGVSL FRDPVIRSSDFEDTPITVNSKLVIASNSLFLHCRPFSVPATVKTPPVTLTNNKQITIN ELANTTQEYDPNAPPTLCSALPPDNKKLRSILKQPPATSESNVQSDCLLADIFFAMGS RQPQIGESPITAFNTVTIMQRANNSIMFLPNLKLKPIQHLFLKHVLLQRLGLENILFH FKMLYANTCKAAGPYQREYFESMLSRVKQRLEDMVFCLNSIESHDFQKDFRVLSRAPQ RLLTATDKYFLMFPPQNRELAIQVGAEVIESICDGTPLSEVLANLSPRVTIQKETGNN

LLKFYALLTV"

SEQ ID NO 42

CDS

CDS

complement (37123..39321)

/note="unknown; ORF 24; similar to Kaposi's sarcoma-associated herpesvirus ORF 24"

SEQ ID NO 43

/translation="MLLQGPVLLPACPATVAANAPSPANSDFKTQLAIFCCLATNNEI
LENVSLEALDRAMQTETTFYACRALRRLVLGEGLYPFIHRQGGIVGKTGNEYAGPGLI
IDDAIGCTFSHIETHTFLPTVFTYELSDTVLVQSDERILRSLYCSPLMVCGVNYQSMF
RILCRYLQIWEFEECFAAFTRTLPEHLIGTCYQNYFKLLEPFKILTLARCPPPCAKLH
LNYLKFNILGFTSDWISHPELHRVQTVIIHNIESNPVLLKNLSKQNKFQDIKVASELI
IDYQNIVNQSLDVNLQVKINKKDPGKKPYKVVVVTPKSTYYLTFPPEVPIFRVAMCMS
VAEHVCHSCDRLYPNTEFLGPGETPRVLEAMFSRIQYAPKDRDYNFIFNADQNPDRYE
QARHDHQTEPLPDMFDPVKHMSLHNFKISVFNTNMVINTKITCWSLAGTFESIIDIPR
LTNNFVMKKFSVKEPSFTVSVFYSDNLCNGAAINVNISGDMLHFMFAMGNLRCFLPVK
HIFPVSIANWNSTLDLHGLENQYIVRRGRRDVFWTTNFPSVVSSKDGCNVSWFKAATA
TISKIYGRPLLKKLSDELNPILSVPYARIDQVKNTIFTTLETRNKAQIQTLHKRFIEC
LVECCSFLRLDLGALNRAARLGTFDFSKRIISHTKSKHECAILGYKKCNLIPKIYVRS
KKIRLDELGRNANFMSFIATTGHAFSNLKPQVIRHTIRRLGLHWRHKAKI"

SEQ ID NO 44

39323..43459

/note="major capsid protein; ORF 25; similar to Kaposi's
sarcoma-associated herpesvirus ORF 25"

SEQ ID NO 45

translation="MEAALEVRPFPYMATEANLLRQMKESAASGLFKSFQLLLGKDAR/ EGGVOFEGLLGVYTNVIQFVKFLETSLAVACVNTEFKDLKRMTDGKIQFKVSVPTIAY GDGRRPTKQKQYIIMKACNKHHIGAEIELSTDDIELLFIDRETPLDYTEYAGAVKTIT ASLOFGVDALERGLVDTVLNVKLRSAPPMFILKTLSDPVYTERGLKKAVKSDMVSMFK SYLMDNSFFLDKSDIAVKGKQYVLSVLSDMVGAVCHETVFKGTNTYLSASGEPIAGVM ETTENVMRKLLNMLGQVDGGMSGPASYANYVVRGENLVTAVTYGRVMRTFDQFMKRIV DRPNAOPSVDDDRDAVADGQDSLAKTPIAAAVIQIGDKLVALESLQRMYNETQFPFPL NRRMHYTYFFPIGLHMPRPQYSTSATIKGVEHPAEQSVETWIVNKNNVLLSFNYQNAL KSICHPRMHNPMPCGQALGQAFPDPGHVHRYGQRSEHPPNMNLYGLVYNYYQGKNVAH VPDVALKATMTTDELLHPTSHETLRLEVHPMFDFFVHQQPGAQAAYRATHRTMVGNIP OPLAPNEFQNSRGLQFDRAAAVAHVLDQSTMEIIQDTAFDTSYPLLCYVIECLIHGQE DKFLINSPLIALTIETYWNNAGKLAFINSFPMLRFICVHLGNGSISKDVYAHYRKVFG ELVVLOOALSKIAGHEVVGRRPASELINCLQDPNLLPPFAYNDVFTNLLRQSSRHPMV LIGDEGYETENDRDTYINVRGKMEDLVGDMVNIYETRNNADHDGRHVLDVGPFNENEQ HMAVLEKLFYYVVLPACTNGHVCGMGVDFDNVALALTYNGPVFADVVNPDDEILDHLE NGTLREMLEASDIHPTVDMIRTLCTSFLTCPFVTQASRVVTQRDPAQLLTTHDDGRYV SOTVLVNGFAAFAIADRSRDVAETMFYPVPFTKLYSDPLVAATLHPLVANYVTRLPAQ RVPVAFNVPPALMAEYEEWHKSPMLAYANTCPMTPTSLSTLASMHMKLSAPGFICHAK HKIHPGFAMTAVRTDEVLAENLLFSARASTSMFLGQPSVMRREVRADAVTFEVNHELA ${\tt SLDMALGYSSTITPAHVAAITSDMGVHCQDMFLMFPGDSYQDRTLNDYVKQKAGCQRF}$ GGPGQIREPVAYVAGVPHSDNIPGLSHGQLATCEIVLTPVTADVTYFQTPNSPRGRAS CVISCDAYNNESAERLLFDHSIPDSAYEYRTTVNPWASQQGSLGDVLYNSTSRQVAVP GMYSPCROFFHKDAILRNNRGLNTLVTEYAARLTGTPATSATDLQYVVVNGTDVFLEQ PCOFLOEAFPTLAASHRSLLDEYMSNKLTHAPVHMGHYMIEEVAPMKRLLKIGNKVAY

SEQ	ID	NO	46	CDS 4349144408 /note="capsid protein; ORF 26; similar to Kaposi's sarcoma-associated herpesvirus ORF 26"
SEQ	ID	ио	47	/translation="MALDKSIVVSVTSRLFADEIANLQSKIGCILPLRDAHRLQNIQA LGLGNLCSRDSAVDFIQAYHYLDKCTLAVLEEVGPNSLRLTRIDPMDNYQIKNAYQPA FHWDNYSELVVIPPVFGRKDATVSLESNGFDVVFPAVVPEPLAQTVLQKLLLYNIYYR VAETTPTDVNLAEVTLYTTNITYMGRNYALDVDPVGSSSAMRMLDDLSIYLCVLSALI PRGCVRLLTSLVRHNKHELVEIFEGVVPPEVQALDLNNVSVADDITRMGALITYLRSL SSIFNLGRRFHVYAFSSDTNTASCWCAYN"
SEQ	ID	ио	48	CDS 4443345242
SEQ	ID	NO	49	/note="unknown; ORF 27; similar to Kaposi's sarcoma-associated herpesvirus ORF 27" /translation="MSIPKIMTVSRDNEGTVCEVAVDNGRHRAMIYYPKTTNLANERA DVVKEAFDTETPVDIVKQIVNEGLAISKKNCVRLALYLYFYLQYVCFALLLTWQLNPY MDPPGLVFAVNPMGPKHVTKLPHPAIVAVGCGADAICKNCSVPDIKTELGMVYHNGSS
				DSGQRAHYGLALLKAAWLVMGNVCPEPVVRQGAALLGPWNRTEWSDFKSAMAATTFCG SRGVLWSPIHEKNLCRPTWNDVINTSVFTNESLCPNIPVVPESVIVLNGDA"
SEQ	TD	MO	E O	CDS 4540845683
SEQ	ענ	NU	50	/note="unknown; ORF 28; similar to Kaposi's
SEQ	ID	ио	51	sarcoma-associated herpesvirus ORF 28" /translation="MTAHTNGVLTTTGFSTSQPESVQVSPFYRVITKPPVMGLFFCVA MCVIALVWYVMRRVCCKGRVVADSCRDPRQPAYEMLNVRLRPHGTNP"
SEQ	TD	MO	52	CDS complement (4573346779)
SEQ	ID	140	J2	/note="unknown: ORF 29b; similar to Kaposi's
SEQ	ID	ио	53	sarcoma-associated herpesvirus ORF 29b" /translation="MLQKDAKLIFISSSNSSDKSTSFLLNLKDAHEKMLNVVNYVCPD /translation="MLQKDAKLIFISSSNSSDKSTSFLLNLKDAHEKMLNVVNYVCPD HKDDFNLQDTVVACPCYRLHIPAYITIDETVRSTTNLFLEGAFSTELMGDAATSAQSM HKIVSDSSLSQLDLCRVKSTSQDIQGAMKPCLHVYIDPAYTNNTDASGTGIGAVIAVN HKVIKCILLGVEHFFLRDLTGTAAYQIASCAAALIRAIVTLHPQITHVNVAVEGNSSQ DAGVAIATVLNEICSVPLSFLHHVDKNTLIRSPIYMLGPEKAKAFESFIYALNSGTFS ASQTVVSHTIKLSFDPVAYLIDQIKAIRCIPLKDGGHTYCAKQKTMSDDVLVAAVMAH YMATNDKFVFKSLE"
SEQ	ID	NO	54	CDS 4690547135
				/note="unknown; ORF 30; similar to Kaposi's
SEQ	ID	ио	55	sarcoma-associated herpesvirus ORF 30" /translation="MENDTPKDKISEADFQQCQAFFHRPIRDLISSGADALNHFSLSE SDGHKLERIVLLLDLVGTECLSYTTIAAKNVK"
SEQ	ID	ио	56	CDS 4709347746 /note="unknown; ORF 31; similar to Kaposi's sarcoma-associated herpesvirus ORF 31"
SEQ	ID	ио	57	/translation="MSLLYHDRCKECQMTRVNSPICRFHNVSNLYQCLDCKRYHVCDG GRNCVIVYTRENLVCDLTGNCVLDNVQDVCSYGPPERRVPDAFIDPLVSHGTRECLKS DILRYFETVGVKSEAYSTVVKNGQLNGIIGRLIDATFNECLPVMSDGEGGRDLAASIY IHIIISIYSTKTVYDNLLFKCTRNKKYDHIVKTIRAQWMRMVSTGDPSRVSATGCFT"
SEQ	ID	ио	58	CDS 4768349077 /note="unknown; ORF 32; similar to Kaposi's sarcoma-associated herpesvirus ORF 32"
SEQ	ID	NO	59	/translation="MDAHGLNRRSVAGQCDGLFHVILPRGFILANNITCGERQRFFAH TWFAASGRTSKTLYVWGRVFQNTDPGRGDGPSGPWSGLAISLPLFTTNGKFHPFDVVI

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LKADTPDSGSSWTVKFLYMSLIAAYRNAMRGLKDKVSQCTDAAVDGEVHPLTVLKEAL VSPDTATRPVSACNPLQMLTGLLQSRVRDDYVTHHRALERPGNVRGQVIAPTRTEMPN GSPSRVRLGFRPPKQANYPKTWAQARHVFSSRAYYVCVYDNEELDTKWQRQDPRPLPL DWSDPVAYLLEGDLFLGAKQNAFVDSIEKTCRCQNYTIKQFFPVLINRDNETVDLIKE HFIEACFVIRNQVSERSAWVKAALFRNDSNTYWKDVLGLWEHGPHKLGTAIKLPTSEP CNADVNWSWLLCDEDITRSISGQSTVCLVVSPTLTAWLVLPGGFVIKGRYDLSSEDLM FVASRYGHPASSHS"

SEO ID NO 60 CDS

49049..50059

/note="unknown; ORF 33; similar to Kaposi's
sarcoma-associated herpesvirus ORF 33"

SEQ ID NO 61

/translation="MATQRRHILKSFLNKECIWLRHPGTSAFVRVYTATTAHSAVFDP
PVTSENAMSLNFLNVMIVIMKPKEFGPCVTVYMNGDILDFCATESVAIRDVPGRADLC
LIRFGTLSNAPRSVPIPGPLNPHPRETVPGLTKQEIIYTSQTVPRGQIPDAIKGKEFH
QINPFLWFDGGAFWQLFLSVDFMLLCPALDTVPSLARIVGLLTQCDKSTCKICTGAHV
HVNPYRGYTPPDSQGTSPSCPCLISCGARRAADVLVTGHVNLLGLLFDPKASPKVTKL
RLKRNPRPVPIEDAMSGVTAEGTEVQPTSLPWALIRLPDLASRVMLYGCQNLKSICLR
SY"

SEQ ID NO 62

CDS

CDS

CDS

complement (49977..50960)

/note="unknown; ORF 29a; similar to Kaposi's
sarcoma-associated herpesvirus ORF 29a"

SEQ ID NO 63

/translation="MLLTSYRERLQNNLRVVTDGGCENWFRQPPVIISGNDKTERMAH PCLGVIHAVNAYSSVLDDYLQTYRRVQEPMPAPTLGKPRISSHATLPRLTEELTNYLK QTCCRVQMANAKDQYMEYQSAQRTHEAFLECPVYAELRQFLANLSSFLNGSYVPGVCC LEPFQQQLIMHTFYFIASIKAPEKTHQLFATFKQHFGLFETTDDVLQTFKQKASVFVI PRRHGKTWIVVAIISVLLSSVENVHVGYVAHQKHVANAVFSEVIATLSRWFPAKNLNI KKENGTIVYASPGRRPSSLMCATCFNKNVSRCFLSSGSRIASRDWLNPAGE"

SEQ ID NO 64 CDS

50959..51942

/note="unknown; ORF 34; similar to Kaposi's
sarcoma-associated herpesvirus ORF 34"

SEQ ID NO 65

/translation="MFPSSFLNNGHPETERRFVKGVQLALDLCDNTPGQFKLVETPLN SFLLVSNVLPESRPVRDCPQPEGFDFEHIHLPKLTRMQRVLGRYCDHVNNDDTCVNVK ASSNSQGALFYLPYGQDEWNWALTLRKDKLVKMAVEGLSNPTTWKGLEPVDPLPLIW LLFYGSRSFCREPECLYERNFGMKGPILLPPHMYAPQKDVMTFVHHVIKYVKFLYVNA GGGLETEPSPPFEASRLRAAIARLGDVEADDAYLSAKCMLCHLYKQNDTISIHETHVG GVIALGGDGARYITSSVRAQRCTSRGDFVLIPLYNIEGLVSMIREHGLGSS"

SEQ ID NO 66

51923..52372

/note="unknown; ORF 35; similar to Kaposi's
sarcoma-associated herpesvirus ORF 35"

SEQ ID NO 67

translation="MASAAAKKMLIKSELESEINKKLSISVFDRFGADSAVFNAQYKG TRESLRSYNSLKKKDDLATVVGTLETSLREKQSELGLLKGFNRKKIEEFDAVADAVRD LKDELYGELEILGTLDNESVPVEEESPKDDIIRWKLERLPRVCPKSP"

SEQ ID NO 68

52278..53585

/note="kinase; ORF 36; similar to Kaposi's
sarcoma-associated herpesvirus ORF 36"

SEQ ID NO 69

/translation="MNLFPWKKSPPRTTLLGGNWSVCPECAPKALDPIPKVQTDVDRT
ASSHITVIKTRKTIAQLKIPNNWGQCSHQATDWTAVLGRGSYGVVRSMSLGRCVKHFG
SRREFFYECIFNDIVRACREKHPLNRGGDRILCFLEPCVPCRALIFPQLTGNLLNADL
KHVNPERLAVEFSELREGVSFLNNICGIVHCDISPENILIKGELTTAYGRLMIGDLGS
ASLHTGTPWTGVMVTSKLGFVQHTYHFKAPARFICKHIYRPSCLLYRCLLSCAGGPQA
HMLNQPFQITPQLGLTIDISSLGYSLLACLEKYLQPADPFPQQGALADASSESAHPLF
YLRCMVPRVVIAEIFSVAWDVPLDLGIDSSGHAPAIPLREAYRRFFANQCSLYRAQYK
EDALENASSRLCNSKLKLVLQKLLVRDYFSHCGNCGDHGFFLR"

SEQ	ID	ио	70	CDS 5356655008
SEO	ID	NO	71	<pre>/note="alkaline exonuclease; ORF 37; similar to Kaposi's sarcoma-associated herpesvirus ORF 37" /translation="MDFFSDEPMVQEMALLDIDEQQRLLSKMSLANFLKHERVRAFFS</pre>
				DNKKEISMPAIRFVYNFYLFAKVGDFIGNTDVYDFYVTCVFRGRRLTRLSEVYDACLN MHPHDRHHVCALIEQVTRGQNINPLWDALRDGIISSSKFHWAIKQQNSSKKIFNPWPI VNNHFVAGPLAFGLRCEEVVKKILATLLHPGEAHCENYGFMQSPLNGVFGVSLDFGIN VRSDPKDGLEFHPDCKIYEIKCRFKYTFSKMECDPIYAAYAKLYQKPSMQTLKGFLYS ISKPAIEFVGEDRLPSESDYLVAYDKEWEVCPRKKRRLTAVHHLVKKCMIHNSTAPSD VYILSDPQETGGQINIKAHLSANLFINVRHPYYYQVLLQSLVVQEYISLSKGTKNLGT QKNFIATGFFRKRQFQDPSCCTIGEFAPLDPHVEIPTLLIVTPVYFPSVAKHQLVKQA TEFWAASAREAFPELPWDLSSLCANAPPTP"
SEQ	ID	NO	72	CDS 5496355172 /note="unknown; ORF 38; similar to Kaposi's sarcoma-associated herpesvirus ORF 38"
SEQ	ID	NO	73	<pre>/translation="MGFILSVCKRPTNTVDVKGEPIDVSKEFDPIIGEESIVLLTADG TAPAALYKPKTKPSKHKNNKLSDFV"</pre>
SEQ	ID	ЮИ	74	CDS complement (5525556391) /note="glycoprotein M; ORF 39; similar to Kaposi's sarcoma-associated herpesvirus ORF 39"
SEQ	ID	NO	75	translation="MKISRSDSFILSSWVKLLVILGLMFIMSAVVPLTATFPGLGFPC YFNTLVNYSALNLTVRSSAKHLTPTLFLEAPEMFVYISWAFLVDGYLLCYYAWAILAI FKAKRVHATTMTSLQTWIVLIGSHSVVFMSILRLWTIQLFIHVLSYKHILLASFVYCI HFCLSFTHVQAMISCNSATWSLRVLEQQIPENSLLDTLLRYGKPIGANLYLSLIAMEM LVFSLGTMMAIGNSFYMLVSDIVFGSINLFFVLTIAWYINTELFLVKYLKHQIGFYVG VFVSYLILLLPVVRYDKVFISASLHKVIAVNISMIPITCILAIILRIIRNDWKWCAKS PEYAPLPQGPKEKTTKVKYSPELNALYETEEDVSDYEDAYPKYI"
SEQ	ID	NO	76	CDS 5652657932
				/note="helicase/primase; ORF 40; similar to Kaposi's
SEQ	ÍD	NO	77	sarcoma-associated herpesvirus ORF 40" /translation="MNAREVALTGHVLHISLHSTHEREKLIIWQVHLLVCQQCGIQGD AAYLFVTETLSNTDWGNIPAINRHAPSINEHGRNYMQWELRTRLRNPIIQLLSRQPGA VNVRVSEPNMVIVGCERALDHSCSVRVTGAYLHCDTTMDFSLDSVVSPTREFWFSEMF SHCLVSNIEVYLKTTGGLYYRASSATQCRKRAKDGALGILDIFNCESREIQVAGQKYT LSIATATFHVLWVDEACMWNGALAEFFRALHNKLFGDREGVAPTLTYVCPGATPEGTP FPPYFSAFPHLPLVFGRPRRLDVTAVQELPKAQIAVHWPPFKDSILGDQLLIPGISPK KPGTVPVRWPLWVEDVNLSLCETTESVARIVDPHSIVIIKFSSLLCQHLKCHRAFVKN ELEYIATICSSDLRLFIQEEYNRLLATIFTWAAASGYTWAAIDKTTVFIKAPQLSAAV SGFCPSLNSCRRKQCYEG"
SEQ	ID	NO	78	CDS 5791758528
				<pre>/note="helicase/primase; ORF 41; similar to Kaposi's sarcoma-associated herpesvirus ORF 41"</pre>
SEQ	ID	NO	79	/translation="MLRRLKITVHFLSQEQQKVVTRLEAHLGLPVQETSHPPDWLKCE VCSASVFLKIPAGVLYAGLARDPTREAKRDSWLDCLVEGATLLLNNSVLPIGALAGIL PTLFANRRCVNFWLLPRAWVKSAPICPPLPIDCVTPPQFVVTKRGPICWYKEWPLPVD VDFMYYLQEALCVFSVVSNGEGTESHADNIRQLEKFEKVLCLF"
SEQ	ID	NO	80	CDS complement (5852559343) /note="unknown; ORF 42; similar to Kaposi's
				sarcoma-associated herpesvirus ORF 42"
SEQ	ID	ио	81	<pre>/translation="MDQILKRLMGEQHRSEAVMPETECSSRGPYNYPVFPRLMLEVHK KNSICMASNTPKLCVRGRLNVPDLGVHVRTRLQSATFTGFVFACVVEHEDMIDALDIY</pre>

CDS

CDS

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> PHVFSDRVQLFKPASASVTELCCILSMLENYDKPPLSFILSALDRARYLHERYTCNDS AFVLYGIEVIASTLAAYHELNPPQGILRVPPLVRFKLHKLLDENADDMKGLLKPIYLE SFRLTENVGEEEGHAETFNIFYCGTIFTRHLHNASVLKYFQITSLHSIPRQTLF"

SEQ ID NO 82 CDS complement (59297..61027)

/note="capsid protein; ORF 43; similar to Kaposi's

sarcoma-associated herpesvirus ORF 43"

SEQ ID NO 83

/translation="MFKMNPGFGSTCLVHPTELSISLFEILQGKYAYVRGQTLHSSLR NPGIFGRQLFIHLYKTALGSCTYDNVLKDWTNFETTLKTRWRGVEHLTPEFKRSTFES WARTVRLTVDQLLLNTINQVLHTRTVLSYERYVDWVVALGLVPIVRRTPDGDTIARIQ AHCQQMRKTYASGDVTISRIVDKLAQEITSIMTDVTSIYIPDYAEVSVEFNGDKAAYL GTYRQKDITVEVVSRPI IYNGRVAFDSPLYRLFTAIMTCHRTAEHAKLCQLLNTAPLK ALVGSTCNDMYKDILARLEOSSOKTDPKRELLNLLIKLAENKTVSGITDVVEDFVTDV SQNIVDKNKLFGTGTESTTQGLRKQVSNTVFKCLTNQINEQFDTISNLEKERDDYVKK IOCIETQLLQSLPEGGRPRHDINILTQNTLQALSGLRDPTINLSECHIPKGSSVVNSF FSQYVPPFMEMLKELTSLWEGEMFQTYNLTPVVDNQGQRTSIAYSQDTVSILLGPFTY 11AKLTHMDLINHSLISLSLHDIADQLYVDSRLSVYINDIGHKYCEQISQPGTDGPNT EASNGGAAPI"

SEQ ID NO 84

60966..63338

/note="helicase/primase; ORF 44; similar to Kaposi's sarcoma-associated herpesvirus ORF 44"

SEQ ID NO 85

/translation="MESSVGWTKHVEPNPGFILNMTSDAKVRGVVDHVSRLSNITTSP PEMGWYDLAFDPAEDSGPFLPFTVYLITGTAGAGKSTSISALYQNLNCLITGATTIAA ONLSRRLKTFCPTIFSAFGFKSRHINIAVRKAHQTGAVSIEQIQQQELSKYWPVIVDI MKEVMAKKPNGMYGTISNANFETLSRMTGPCLWTSNIIVIDEAGTLSSYILTTVVFFY WFLNSWLNTPLYRQGAVPCIVCVGSPTQTNAFQSTYNHGTQKTEISSCENILTFMIGK KVVSEYVHLERNWALFINNKRCTDLQFGHLLKILEYNLPIPDEVMSYVDRFVVPKSKI MDPLEYIGWTRLFLSHSEVKAYLTNLHTCLTLGGDTRDTKLFTCPVVCEVFVKPFEEY KRAVNLTNLTVTEWVTKNLFKLSNYSQFVDQDMSIVATESTERSTQVTFITKFVKNSH VSLNGKTKKCICGFQGTYFEFKRILDSELFVETHSQDRPEYVYGFLNTLLYNAMYSFH AYGVTRSHEKYLQDLKFAPLPAALATGRVDLQTVREELNLEDDIFYHVCSPPPPAGIT SLQVLVDTYCALKDVFASRIKVACRWFGGEFEKETFSAFTVNMVVRDGVDFVSPSERL NGLLAFASTVESYKIKGYTFLPVAFGRCQGLPLSDDLRKKMPSLVVQDSSGFIACLEN NITKLTETMEDGSVFOVCCAGDYGVSSNLAMTIVKAOGMSLERVAVVFGSHKNVOTSH VYVAISRAVNSNYLVMDSNPLKTLLREPVDNTSAKHIVRALHNPNTTLIY"

SEQ ID NO 86

complement (63379..64437)

/note="unknown; ORF 45; similar to Kaposi's sarcoma-associated herpesvirus ORF 45"

SEQ ID NO 87

/translation="MAMFLSDPSRTPPATPRMLPIPGAPRKKRTRFLFAGSRTGLPV PPGYGGPPVIDMTAPEDVFDQDSPPTTPKTPDETDSHSENSDYSDVDEEDEPPVSSPP RIDPHARDGESFNOSGRLPTVITSTGATTPPSAPAPLTAFGGPRPVAVVTGOHRAPOS SESDSEDDFFIDDYEDTDESGGEADGFSPRASPAWSGDISRSPAEGGWSSDEEEPVVA GSTAGQETIIISDDDEVDDRGSVETWDESDADEGTGATDVIDLCSSSDSDDDADHVTS GGVRAACKRRASRRDCNGDDDVIYVGTTQGPKRRMTSTTGGGATSNPEGPGVSGRQTM AATPPVCGNDNYPWPWLD"

SEQ ID NO 88 CDS complement (64479..65246)

/note="uracil DNA glucosidase; ORF 46; similar to Kaposi's sarcoma-associated herpesvirus ORF 46"

SEQ ID NO 89

/translation="MEGWLKTIVWSKMSPEVLEEPSTQTLLLSDSWLEFLNLSPFLKQ KLAALLKRVMDMSNVTVIYPPIDRIMWWSYCCEPEDIKVVILGQDPYHRGQATGLAFS VAPDYSIPPSLKNIFKEIANTVPGFTAPSHGCLDCWAKRGVLLLNTILTVERGKAGSH ANLGWDWFTSYIISCLSAKLORCVFMLWGRKAIDKATLINGORHLVLKARHPSPLATA HAATGSPWPOFLGCNHFKLANDYLVONRRGAVDWNIN"

SEQ	ID	МО	90	CDS complement (6522265731)
				<pre>/note="glycoprotein L; ORF 47; similar to Kaposi's sarcoma-associated herpesvirus ORF 47"</pre>
SEQ	ID	МО	91	<pre>/translation="MRSMYTLSLFITCGFFLITCCTGLVVNPCCKIIPLSDFIFPEPF EIASFHLTNLALCPGLCTATLRYKADRSTTEICVNGFHLRAFFIRILYKLNYSVPREE LQLLNYMQYSLDEFLAEFEDFHINGSESGTAYTRPPLLDFSDRSTKVSRIRKVITRRG DLWRVGLKQ"</pre>
SEQ	ID	NO	92	CDS complement (6599967168)
				<pre>/note="unknown; ORF 48; similar to Kaposi's sarcoma-associated herpesvirus ORF 48"</pre>
SEQ	ID	ио	93	/translation="MAVSIPVKGVNRETESNWRSIVTTFERHGNADRAIRSLLRFFKG VDHPGFLASLVILKDVTIDSEKTIERTDLIPLLQGVRFVTQQIYMHLKDHASESPMAE IWRDCKERFCLALELACGCQRCASAARQLRACQQACRPPKLNPHKQQCVAARLLTAVY NQMVLRTRVSVSEFCLNALMCVPREFGFVSGDVRVETSRVASCLNLSWLYLILDSYVR TDLTNLEMAMSRACRIHGLSAGDPFYSALVWLKNSYACDTNTFFFTVNSTSVTTPILM DICASLTGPVPDVIKINMLPLVNDQMHPSVCVERANFTGSCPKVSPTHHLDGLKLETT SLTLAADSLDDILQALELICGDDEGILDSYISDINTETEVDESSIEEEIVFEELS"
SEQ	ID	мо	94	CDS complement (6739868303) /note="unknown; ORF 49; similar to Kaposi's
				sarcoma-associated herpesvirus ORF 49"
SEQ	ID	NO	95	/translation="MSRHYGKDHLLNHMYKFHYPPLGMIVGEMNTLTVNARNPLYQAA TLRVERALYLSKILQVLMQHRQGERFIVPQCRSNMVYCLKELHKITNDRIRGLINSVL PLVDAGCVGFDEELVRVLPEILKLEYPHAHELLPPHDPTSPLSWCLSHMVGVTKTFKG EVKEMIDTFHDLSVPSFQYLASLVKKFFLVEEVIYEDYQDTQFNVFLNLCFFWTTVIK MYQSCIFKDKLLDTIKACIELLKGEARQFFGWYDLNTPNLGSSALVKYTEHLIRALSV DSSAIPIGEICSHLHHCKHALLNLE"
SEQ	ID	NO	96	CDS 6849470038
_		-		<pre>/note="transactivator; ORF 50; similar to Kaposi's sarcoma-associated herpesvirus ORF 50"</pre>
SEQ	ID	NO	97	/translation="MECASLGPISGLIADLNLFNLFCLYRGSRVKTRGAATCNVPCAE CAQGIVRILTERALCCTEKMFIASACSGVVIPPQLARVLHDVYAEMKAKCLGAWRRLI CCRRPIMAIADSVLVTYNTLDAEGKLELRLKALCKLVFQPIFLQRILAPMQLLANGKM VPDNYFTITGTAEKRRPVVTGSTSGMTCPGSSLVPDSLILPVCEPGLLPAPLVDLSNV LENPEIILSAPPLSQFVITNTHPSLPQSVSIITPTQGVVPGQCFMDTWKAVSQSIHHQ AQTPILAAALTGSTSAAPGPHIACSPVAGTSRQVEGSAGVDCGKPACVPQPALPPNVP AKRMETVAQLGNAPVKNVHIGGRVYAPLVNIPIIDLTSPSGSGQSPADIANTPESRMA AGSPPFAETAATVPAKRKQPREDVADKRLKGDVRGAATVNHPFPGPSGMRVREQGLFD
				LIESSTDVTANASGPKNDDDMLAAILQDLYGLQSPPAIDSPSSNSDNEEIFPEVSPPS SGHGSP"
SEQ	ID	NO	98	CDS 7035570888 /note="R4"
SEQ	ID	ЙO	99	<pre>/translation="MPRVKTQPKRPQVLEFMPLDLHGGTHTEMDSQNLCPDGQDLLGS YIYTENNGPFSQIMHNGQSNTGTGESFGSYAAGDGFLGGSVSGMYGNNTGEGACSKRP SACRKRSAALIHAASEASVAEQGTSQGAHAVSDRIGRDGGADNRLLKVSARLSDKTKS ALRSHPCLRCYSLMFNT"</pre>
SEQ	ID	NO	100	CDS 7146872160
SEQ				<pre>/note="R5" /translation="MGFGNIRLGWRLCFMVWVAXIARGRSVCPTWHLTDGKYEAVYRH</pre>

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VVLSFRAIRARSTRDTEQSVRDRNTVTTSYRTPGRPSLFQARPSSHGARLPPSPRTMA RYAESRTICDQN"

SEQ ID NO 102 CDS complement (72401..72820)

/note="unknown; ORF 52; similar to Kaposi's

sarcoma-associated herpesvirus ORF 52"

SEQ ID NO 103 /translation="MSSTRPKTRAPKKELTMEELAAQVQKLSVENKQLKKLINSGDPT

RSGSDPVISNTEKEAKIAAAVSALCNVATRKIEAKVRAATAKAVTRGQMEDALAGISI

RVDVSMDETTRGGIAASADGALRRRRAQSRTRNNDAD"

SEO ID NO 104 CDS complement (72884..73198)

/note="unknown; ORF 53; similar to Kaposi's

sarcoma-associated herpesvirus ORF 53"

SEQ ID NO 105 /translation="MTGSIVLALALLACLYLCLPVCATVTTSSTTGTGTPPVTTNPSA APSVTPSFYDYDCSADTYOPVLSSFSSIWAVINSVLVAVATFLYLTYMCFFKFVETVA

HE"

SEQ ID NO 106 CDS 73274..74146

/note="dUTPase; ORF 54; similar to Kaposi's

sarcoma-associated herpesvirus ORF 54"

SEQ ID NO 107 /translation="MAEVTAHTVPYAFDSCKFEIIPKNNSSRIALRNKFPVVVKPGEP LVVPLGLKIIRAPQCAFFLSGAPTDEVYYHTGLIDQGYRGEIKLIVLNKTKQVVTLYR

GEVNVSLIAFMYASPGPLKCPILNLPHYSLDAGFDVTSPHAMTIPPTDRTPFTLSLYY KSPQLSTPHVPLIVGRSGLATKGLTVDATKWTQSLVHLRFYNFTKEPIDIPANSRICQ VVFIHEDHVPSGWNILRSRVOLGSTLQISWAKIRFTDVATLPKTHPLNSRHTQSQTEP

ETARGAKGLGSSGL"

SEQ ID NO 108 CDS complement (74207..74839)

/note="unknown; ORF 55; similar to Kaposi's

sarcoma-associated herpesvirus ORF 55"

SEQ ID NO 109 /translation="MAAPGSFWTCCGFSPFGRVGCQYRPLPDPLNECPTHWRTEIAMG LPPGVDMGDVKQAEMCTAALRQTYLLAVQSNKITEYLRRFDAARVPAGCQETVRIQIS

LPPGVDMGDVKQAEMCTAALRQTYLLAVQSNKITEYLKRFDAARVPAGCQEIVRIQIS KLKSIQNVIWNAMLSLAIGDITVDESAFHALLNKRADETVSLLEMEKLATTIASDDSV

TWAAEINNVLVDTEASSNPSHPVIRQPTPQLAVADNIVPDKIIQDAQADG"

SEQ ID NO 110 CDS 74851..77337

/note="DNA replication protein; ORF 56; similar to

Kaposi's sarcoma-associated herpesvirus ORF 56"

SEQ ID NO 111 /translation="MVDEIRAIFSTSGDMAEVITDILTETQATASFFCVLHDRGDAPI NTPHAVIKLCLPAKRPGGGPRCLPLMVLNLPAWQVNLFLTGDAPLTSDNIKDRIDLAQ

TEEILEPILSVLACKRSAQQTKHDSFKSKVAWFRAKFVSALRKVYKMTPSPYWMITLL GSFEASFVLAGTFYFFQSHICTAETLVHLTRLFSSSQGQSLVTVNTYDELGRVFGRSD FLGIVPNFWAYLKYKMQQDDVESRAIDQTINSIRGGLMLSPQDLVHFIYLSFYECMNA QTFLSYSRTTSSLPTPATVNPPQLCRRLEADFKEHVMAYYNKASYLSTYITILTVPAP LPDGYENFQELACQYWCGQSRDVAEIMTRINDQYPQLNLTKDLSGLLDLAALDQYSGG PKENLFTVASRIPTYRCEFLNKQYFVLMHADCIDAYWKQNIIVPEDAQLQGLTDQDLT SRIFYCDLGLSLPTFKQQILVSRHEYFNPRLPVYRWVLDFDLKVTEGRRTLNDIYNIC

SRIFYCDLGLSLPTFKQQILVSRHEYFNPRLPVYRWVLDFDLKVTEGRRTLNDIYNIC VTLRQVILETLQLIGPLKPNHPVYFFKSACPAVTWPDDISDTAFCHCDAKIGMRIVTP FPSGYCLVGSAPLVSLTDILNRVVKLDTRLASEYPGILEDKGPFDSGIYAKGRCVRVP

HCYKVGPGGELSRLLKIIICHPEESDKSAYLKNAFKVSNLLHHAPGDSVTKNGHLVYA ITDENEGFLESKTKNNLPKTITDLAEKIERTTEKPLIDWAATAVWPKLHDTIQRFFPD DRIGQFASVSFMHSGDNIIQVKPQKGNNFFCINHKHRNHTQTVRVFLTLHSTKESEVT

VTFMSQCFAAKCNHNSPTAHFSFMVPITGT"

SEQ ID NO 112 CDS 77578..78906

/note="immediate-early gene product; ORF 57; similar to

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	Vanosi 1s	sarcoma-associated	herpesvirus	ORF 57"

Kaposi's sarcoma-associated herpesvirus ORF 57 /translation="MRYVFHALICFIGGISSSDFDDSSSDEMDDLSPTPEPEPSTTPN SEO ID NO 113 SFPEGPKSQVVALPKIRKRSRSETPVKIEHRSPLNRSRSRSRTRSGSGQRSNQSGRYV KRFKPTVDAPRHREPWHRGGKGKAPFIRRDAMAGRGRRTYGHDYRGKAALTRSIKESI KKMHLPSTMLSRAHDKKVFEGLLPRHLGQCFQVCLPAPPPLQPEVFTDRQLTAIVKSG GRRDALVAKKVSLAKLTSLYKPLLTFVTGRNNQAHWLATRKNTLASAGLEALAAFIEE GLAWAQVCVSQNRSLNDSNLDIILDSSQSVCTWFISKIRHLHIQCFLENQGEVSLVKQ LTYLVCINNRLAEAANLAGEVKLNFKLGMLIGFALTLPALLAEHKLSGESLYLFRSFL EKYRPGDVMGLLNSIVVEHYTKCRSAECVITTHAMVGSGENNKGLFFFPV" complement (79266..80513) CDS SEQ ID NO 114 /note="R6; similar to Kaposi's sarcoma-associated herpesvirus vIRF K9" translation="MATWRPPQSGGPSAMGLREWIVTHANLATYSGLFWADDEKTRVV/ SEQ ID NO 115 LATTTPWSVGFDYLRDGKMYEDYCNQRNIPLPSGRSRLGQAKARLLGAIRKSAYFIEE KDVLRRSFSFANVVFRLRSDEEMLCRLCPRASGVAAELRGLRFRMFKRKGADEAGRVS EYTVKQLLGLLRTRPAGTFTMTAPATEASATATASGEDGRQDNSQGGAVALPGEHALP LSASSGLSACLAPSVDDPWGFMHIQVYYYGVLQAQTFTHSGMGVRLSTRPTDKNEHHV CMAPGPLQLWLPPAPYMDDDFMLSRLVNALHALEDGIVLCSCQYGIMMNGYGFLNLWF RGNTSNTSEPRRVPSGVGHRVFDTDEYMLKLAQSPRPSDPGPPDPFAQIWVSAWSLYE EEDOSOAPICIVVHQREIYRHFE" complement (80686..81933) CDS SEQ ID NO 116 /note="R7; similar to Kaposi's sarcoma-associated herpesvirus vIRF K9" /translation="MAGRGVDIRAWLVAAVESGEYRGLVWENEDKTVVRVPWNKVTAD SEO ID NO 117 RSVWNSEKFFDDYCNMRGICQGEKPSHYGRFRKMRFLYDMRHHKSIRELKFINKAYGR PGARYRLFRLLPEPVVSCAMCNLMSSTETQCLGLISEFQYDQGGGSGRERRRVFTATV LARSRMDKNKRVREHRLPGAIQLTFLYFGETVGLERVHAGIRVCSRPCPVLAGHACCF QDERTLFLPSPGVVDCQFAREDLRVMHKKCEKGVLITLTDTGICVKNLENREMKVLTN NEEEYKDLPSRQPVQVFDMVDYLRALARSPNPGDEPPRDYAQIALCLSVQSPNPVDAP IAMRLRYVCETSSVCGTEGCFYPGTTVTSEGRTDCSFQMEDPGEGTSQSHDPAVELGD SGPDSMDDPDAGTSGEDDGVACS" complement (82262..83317) CDS SEQ ID NO 118 /note="R8; similar to Kaposi's sarcoma-associated herpesvirus vIRF K9" /translation="MERPVRVTKPSSLRGWLVECCETGRHPGMRWVDEERTLIRIPWN SEQ ID NO 119 HDRGSRGVEEAEKNIFIDYCRSRGILHAAGRELTAKECKNWLSSAIRHSQTVSDVSTK DNLSTPYPDRCRIIRLLPITVRSCARCDQASGTTAMLRGLREEAVNKFGPVGAGVRYT GAVGAGGEQCWMLRIMFYYYGDRVGEVVTESSNGIRVLPLSERRPQGHICAAPIAEQA LVPEIPGHLAEFQAEALRFLDKDLLRGLAFWADPSGIYIRWLGHSLAFVQGNVESTGA VAVLSCANACRAFNLVDYMTAMARTSPDGAALPQACVYLYFGGVPTPEGGVQSTVPLI IOLWHECLWRALSAANV" complement (83491..84252) CDS SEQ ID NO 120 /note="R9; vIRF" /translation="MDSGCYACILDENSEGIINYLEQVCGIGLEPGMPLPAPLPTLVP SEQ ID NO 121 PTRSAYARAHRLGVPEAPLPHQIVPFWRLRIQVFYFGVLALDHTSQDRRGVRLHPRPV PHPGHLCFYGTGFTVWFPSPDREKLTAEQITQIKTMLVAYNEGIYVHGNETGVYVDNR NRETLYAAGNDCNGDIIQREVMFLSKQKIFNFMGFMRKLARSPGPESHAPCNGATLYL SQQPGAQESPQVPISVVVCQDELVQGHMNPSRWCS" complement (85052..86209) CDS SEQ ID NO 122 /note="R10; similar to Kaposi's sarcoma-associated herpesvirus vIRF K9" /translation="MAAGESRRGPSRYGMALKEWLTFKADSGLYPGLFWADEQKTRLV SEQ ID NO 123

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LAATPPSFPNYDYORDGOHYDAYCELRHIPLPSGRSRLCOARGRLLGAVRKSKYFEED KEFPTDOFPFTALVFRLRSSEEMSCPVCPRVCALRLELRNMRFAMLGRGMLHALSGPS VSDQERRYREGHQDGHDAQDDDAAYSSGLLRARLMACAAPSAGDPWGHMHIKIYYYGQ LOAELLTATGOGIRLSSKPTNKAGHHVCVLDGPLQAWFPPIPQTTESSVVQRLEDALK WLVDGIIFCSTSRGIMFTITGGPNVWFQGNTVEPYSLPHRAYTGMHVWAFDTDRYLLD MARSPSPRDTGPPAAFVKLWVSGCSLGEERNSSRAPLSIIVYQTEIYRHFE"

SEO ID NO 124 %

CDS

complement (86355..87527)

/note="R11; similar to Kaposi's sarcoma-associated

herpesvirus vIRF K9"

SEQ ID NO 125

/translation="MAERDMDLKAWFIEAVESKRYPGVEWDDEDKTIIRVPWNRCTDS RVDEDYNKIFDDFCSARGVCQTGSHAQKFKKIRMLYAVRSHRYLRELTPPSKAGGVSG ERYRLFQLLPEVTVGCDLCNLIATTSLHSCSMGSCVREDVFERTRRPRAKAPLRVSVY KRKSKRLQDSSAQPVLGAVEVSFFYFGENVGVQILRAGSGVRICGLPDPKRPGHLCCA DNPLTCFLPSSQLIPCEFARADLQALQKTCERGLICVMTESGICVKNLEERNMTALTN YSENYYELRPSQPLQAFDLLHYLRELARSPTPGDVPPRDCAWIFMCPSTQSENTWDAP IALKLRYVCNDDVSDDVSNGAAGDDSGDEGPSGAGVGASGTTGSTSVSTLAPYGRK"

SEQ ID NO 126

CDS

complement (87894..88961)

/note="R12; vIRF"

SEQ ID NO 127

/translation="MAEGRAGSIRVNRPSGLRAWLLDCCDNDKHPGMHWLDEEKTLVR LPWNHLKGAGGVSDDERNMYLDYCQFKGIRQTGNRRLSVRECKNWLASAIRHSQTVED VSTEENLSAPAPNRCRVIRLLPIFVRSCPLCNEADATGGMLLDVRNEVTARFRYLGAG MEYEGAVGGDGEQCWMLRLVVYYYGRLVGNMEVGSPNGVRLLPAPKRPLQGHVCAGIR PEQALLPHTPQDMFPHQTSMLKWLGKEIIRGLMIYADGSGIYIRYMGHVPAFLLGNGG SLEPVDIINNARVLRVFSLAQYLSAVSATPPHGTRFPAAYASLHLGGVPTPEGEPCPT IPLSIQIWHECLWRACGDAAQ"

SEQ ID NO 128

CDS

complement (89122..90216)

/note="R13; vIRF"

/translation="MTEJEITHNHLRRWIISNLEANTFPEHLCWCDEEKRSFRISWHR SEQ ID NO 129 GMSGMQPVVAYCLDRDLECGRQHNVSECRKRLLRVLRENAGFEQDDARATTTRFGGER FFYLRPAVDPLCYACILDSHSETVLNYLEAACVHGLEPGTPLPPPAPAEADGAARSVY

ARAARLATVAPPHPDQITPFWRLRIRVFYFGSLVAEHTSQDRRGVRLHKRQDPKPGHE CFYGTAYKMWLPKPOLDGPLTPEORETVCEIINGCEEGVFLHGNELGMYVDNRTRHTV RCAGNDAEGNHAQRAVRSSVKSQIFYVMGLLRRLARSPVPGDTVPSNAVTLYLGGRPG

SSKRPOVPVTLVICQDELTHGDIRAARWIL"

SEQ ID NO 130

CDS

complement (90462..91544)

/note="unknown; ORF 58; similar to Kaposi's

sarcoma-associated herpesvirus ORF 58"

SEQ ID NO 131

/translation="MGTYTSEASLAWLSFMSGTVSASPFILCFIYHSLYFVEPLISVE NIIFSWGAVGLHGLLLLFCIFGLPAWLSRQVDVPCTISAFLITAGSMASTLGVDLPWV HVSIFVGSCLCLLLCVVAANDVVYLCPTIAHRYYELGFLAAFSVYYFLVLKNLFLAPV FLLPLVAFIVGGVCSLRALRSHPLYEAGLQRRHAIFSLTSRRYITYSIKQALEVCGWD FYLVTVLIGGAAAGTLSVGLTTPLLLGLVHYFFVFHVGLFCCLGLVFRSNVLALVYVL AAAVLLTLTHVLGPGTHNLFTRVCVFTVFLLTMFGAIGCELQIIRKKLQRAANSPRIV

LGVCACGNLLMAVVFFSLNKVELGAL"

SEQ ID NO 132

CDS

complement (91555..92739)

/note="DNA replication protein; ORF 59; similar to

Kaposi's sarcoma-associated herpesvirus ORF 59"

SEQ ID NO 133

/translation="MPVSFHYGARVDVDALGSISRVYDHIKGIVKKGVIQISGQGRAP VLSVLSSVGDAGVLGLRLKNALAPLMVYSDMTDEVSFSFRNTSLGNTFTHTREMFGVN IAEMNVAFYHHGDESDAEGKPQFVRTTIAYGDNHTSTVHKSVVDEPNLPSFHDRLEQA GTGNRLFLTVKTLTLLLKWLROOKTRAKOVVTVSLSETLAVATFTVDGVSKIIDFKPD TPDAKWTCARGRKLDVGVVSSDLTTHVSLESLVAALNACKIPGFFLPGFRWHANEILE

CDS

CDS

CDS

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VEGLPLTDSLADVRLGVMLLKVDPTDRNNAVPGNLSEGADPEGVPELPSPPRTPDLDL KEQCVPIAEDGAEPTDGGAKSLRTSGSRPEKKHGKRKHSSSPSRGKGKTKTPRATFNP LF"

SEO ID NO 134

complement (92868..93812)

/note="small ribonucleotide reductase; ORF 60; similar to Kaposi's sarcoma-associated herpesvirus ORF 60"

SEO ID NO 135

/translation="MFGLSIVTAAMESPDRFLYASDHPGFLALTQETWQNRWFPSQIS LHEDSDEVRLLSPTDREFYQFLFTFLGMAESLVNFNIEDLVKEFSNHDVTHYYAEQVA MENIHGKVYANILNLFFGGNRGDLMIYAKKIVEDATLAKKIDWLHSRVRKATTRAEKV LLFLVIEGIYFISSFYSIGLFRLRGIMRGVCLANDYISRDELLHTRAASLLYNTMISR DESPSVAYIHGLFREAVEIETLFIRSKSRDVTMVNVGDIEQFLQATADRILKSINIPP

LFGARPPNACPLSYTSAKSVNFFERDNSEYVTSVHNDL"

SEQ ID NO 136

complement (93794..96160)

/note="large ribonucleotide reductase; ORF 61; similar to Kaposi's sarcoma-associated herpesvirus ORF 61"

SEQ ID NO 137

translation="MNTETSFSAAKSAKPLTLVTDAGTGGCSSSLDPERCAESLVNSL KATLGWDIEANSLTGLLWHRIMEDRCLVTVRDYLAVFGERLSDEVRAFMSKHEAALDG LLQDFKQSKAYTNFVNCGYLSAVRFYDTYVLRTQGSSPIFESVAQMFMRVAVFVACQC IKFPCLRETLRHLVESETELDEMYLVGYAFHYISSQIVCCATPVLRSAGLRGGQLSSC FILKPSMATEDKTLKALHEEMSPLLASKSGVGIDVSSFAEHKNITSCLKLINAHVGYF NDNNIRPVGASAYMELWHHQICDFLNAKMPENQERCHNLFQGVCVPELFFRLYETNPD GQWHLFAPEVAPNLLKLYGAEFEIEYNRLVAAGKHSSSLPLKSMMYALINTVIKTGSP YVLLKEALNKHHWCETQGSAINCSNLCAEIVQQPEGQASVCNLANISLPKCLRPHRGE SGVEPGKGDVTFGFELLDDAVEAAVIIVNACILGGTAPTESVRRGQEERSMGIGVQGL ADVFAELGFGYLDAESAKLDVEIFQAMYFTAVQTSHEIVLLGEGTPFRGWERSRLAQG VFHWQTWDGVKPSHPPLERWEQLGRSIAQHGIFNSQFLALMPTAGTSQLTGYTEAFYP FFANIASKVTSKEEILKPNVTFFKRVKPGDLRTVRRYGGDVASFPEPLKDRYKIFLTA FDYCPIKQLERAGARAPFVDQSQSLNFFLKEEQATRASYIRDLLLTGYRLGLKTMLYY CRIQKQTKLNALQCLDQVVGDNISSEGAESNCVQKADGERTKVCLACQ"

SEO ID NO 138

complement (96163..97158) CDS

/note="assembly/DNA maturation protein; ORF 62; similar to Kaposi's sarcoma-associated herpesvirus ORF 62"

SEQ ID NO 139

translation="MKTRDANVNKLNDSLMRLLPPPPHRVSLSRGRDFSKGVRDLLSK/ YVVSTTTGVEAIKDGFLSVSPTCQTYGDFLIYSQTMSSQEPRGTYLFSFKQTDTGSSI DMLFTPTSLARLSRMDADSAPQTNRIACVWYGHESGLLDAIPNFEELLETGSLHQFLA PVGPLVQTVHSTFVTKVTSALKGNVVAREPVVTHIGLTLPSDMFVDLDDSCPSSLRDE PLPAHSSIYVCLTYIRVNNRPALGLGFFKSGKGYCEIAAQLRDFYSGVIRTKYIQLQN DLYINRLAFGVVCRLGSVPSGLQPSFQSLHFKGAALPVLKFTEFVSNPGSWKLFL"

SEQ ID NO 140

97157..99976

/note="tegument protein; ORF 63; similar to Kaposi's sarcoma-associated herpesvirus ORF 63"

SEQ ID NO 141

/translation="MASSIPAARADNGDENTGGLYKLTDNLLTCTGSLQQLKLLMEFQ LKPLPTAHLLSMPTVTRFLNTAFKIDNPLVSFIQKHPVFFLMRVARLPEPVITDHQSA ETSTGILSEVVNVLNTAIRKPHESPAAKDNDYLDNRAILAMITEYIHHVTSRTPSGIP PTPPMGISHLPCVEQILHETHRQYWNLTLPESLFIDIGEVASPLQTWLILSYCKKLQL APPPLFPPVDELARRLVTGHHELFVPLSTSLETYITMPVSKQRAFEIYSVFAKSKNIV DGTPILAFTDTELTTFTPELLFLYDFVIESLCKNQAYGCSRNAIEHFIKKGIDFMAEL GAFIEKTCGYRSTVSLSNVRAVKARLASCGLSKEACEDFRAMILMTPHETTPKWENFT DFLEMVNQLTLYGFYFYECLNQYSPTSISLAKIQNILNRVDAEQSDRALWRTPLIGSF PFPWKLNNVLAFFKPSTPVATLQKIYKAIPSYLMRSLFEIAANKSWGNIALAESAPLT DIQTAEPDQGPVSAQVIAKYCSRLQISATDYDAAIVSSPGFAAEFIKTKLYPILSEVL RNTSKKNRSLFQIRWLIVFAAEDARDLAPIRRSLALAYFQIMDILEEKHSPESFYNLL DYLQETFRCIRQVIPEATCPQEFLQYLFTFQNIPIAASFIQTSMTFVDDLKNGIPGIL

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DLVSLGAAFYNMKLLYDSTLDTVEIPTEEGQPIVVSMFVFKSTIRVLEKLLQEAVIAL TQTSEPMYAAHIRLMQHLTYMQKIAGHEIMTTQLPSVFHEIHEGYLQCFKRFKRLMLH VTGSCCYSLTRYFGFLYQPPLIPDTIVQKILNFNDKTDTTDDILKSLSQPVRQGPLSA ENESSSRLSKNNVELLQKLYDDFRTASTNNNPTSIKLEYSGNYNETQVSVDWSTYNLV TYTAPDDTLKFTPVNTEALDRMFAE"

SEQ ID NO 142

CDS

99980..107626

/note="tegument protein; ORF 64; similar to Kaposi's
sarcoma-associated herpesvirus ORF 64"

SEO ID NO 143

/translation="MELPPIFSKFKIEGVATTHOADCRFGOYAGSOCLSNCVIYLAOS YFNRESPVTDTNDLDDVLROGATLDFILRRSGTLGYNOYAOLHHIPSFIKTNEWTAAI FQSQEYFGLIGLDAAIREPFIESLKSILTRNYAGTVQYFLFICGDKAGAVIIKNKTFY LFDPHCVPHVPNSPAHVISSSDPTAILEYVSPPDREYTGSFLYIMPSEYVNPEHYITN HYRTITFAKVHGPHIDISTGIEPCTIEDIPSPPRSPDVTSKSSNLARVPRTTTDTSSA KPPPATLSGLRGAEPPTSYPDPATNDADTKLLTPAPAQTAVDHPEFQTTPGATLLLSE LSASRGRKRKLSSLQRYSDSDEASSDDEGAPRRRVHDDAISAEVIWMDDDISPLYSPS ATPSFDDVFDSPPMSPEFTYEDATEDTDGAFLEQIARDAETPFSAFDDLITDHDFSSL DKKIEQLIKYEAPSQHLPNISDKQNGRAVREAAALQAMDKIMINIILEHGLITDAQAR GPSACKNVLQFFILWGEKLNIPISDAKQVLELDLQLIPLHTAISEGKFKQGAFKKHLT TKINRCLASMRATHADAQKKLASAFNVEGSQISSSEAKISVRALKEQIANHLSPGFLA VYSADEVKHLRDKIQDLKTGIEQRNKEIQQEELFFDAMLTALDTFQPPPKTAFPMEIF PHRKTEVMLDHLASITTRLTEDATEALNNYLETPPDQGTHITNIPNFSSIVANIISTL KILTYAENDMQLNVTPMATYRRQLLYLGGELATIFNLEWPYETVPPVQELPLVARAKA KMESVTKMEKNQQALDQILGDAETLLDTITATSGDENPVRAMSIPILETYITNAGALI GSSRNQRFEKLKAAIHDLASSESFIIMLLNNTRLDNISDNLAKIDGILTNNTRFLSNA TVSKTLOTLGGSLIRECVEALNKRSPSSLNNARLLAVOTILGHASVPDHETLTRIVSG VASAQKESAGDDPDRWTRVTGHLNELKLVTTQSRVDKATRRKLLMIITRDLKEAEVSQ ETVLETRWQENVLKFQPSTSKEIEDFLQSAPSAKARKFAEKHLRTLITQFNGHERPPS EATAVPMDYTPTPIPTPQAVSTATAEKGKAAWNKIQQAFQDFNFHLIDASDWQEMASE YSRHGSSLPGTVGPKLVRFMESISNTLDDILTQKLASLLPNGPAFRPPAFDWIAPYQT RVNAFLKTIGLPMVRNLADKIHHQCQTVSHAVQSADLQQATVGTSLERPAAEYCRILS DMQVAFNDHGIAVRSEAAAYTDAINSPANVVTPPKPNLEAPKKLITATDALTVEDFPD FLKTSILQQEQRLIALQRAEFQQLEASISAAERLRQSTRDEIAGKMATAITQLLPRAP VAISSRPLNLSKPIDFLSSTVYDKILDKEPYETAIAGFAWLEIATKSVMVYSQQNETQ QLNVLLSEVEKQSTVAQRLHDLELSAKNTDDVKVLKQALDELAPLRVKGGKTTVDAWK QKLES IESLLRATRTAGEISSELER IGTQAVGT ITVRDLGTLSDQCREAANFLRQASL PEGFSDIGTKLSELQAYIKYKKQFLEHFETTQPNVFQRFPLSQNITENVPARPAMDSV ARLTNHLHVRGSAPHFTTWIETLPTVDPEKPTHVPAHGGAPLHRQITYSNVLEALFSL CSTTLTPVPTAPGLEIATRARRGAEAATWMDRQWPDIAQTLQDVLDTYEHTTAHANRD AAFNTFLAMCVFTOIIRGASRAVTLPKLPSTAVDFPEEIVLTPRECTTLVTAMWPTLA AAILRLKSYSEALGLMSRFLPLMFQALPHLTLEAQVKNGPHNTPPQLRCFAKTEAIPY FPAQWQSANLEQSLWGQTDFLQICDNNQRKARVAAVTWALTTIDGVVLDQLWSTFKPM TAASDDTYVDLVETLHLTTFGPRGPTPRRETTTEHPPYEYGQPTGYCISGQSTTPVQA SNTPVSAFEAVLGAMVFHVPIRIFLAATPKRLGQARGGMGLLTPILECVPDVEPFKSL YNAPRKPVPIETLPASLHPHDERQVFLRQAQWLSYRFTPHEAARSSTPPLLVVIDPEN LVTATYSSGGPANFESRPFYVMPGPYPPDWPKTLSVTSNTSVTHLSHDEICNLFTTLS REHGTVOGRDIFAAAPTNVTPEQTANPPAWETDNRLITQTETAKKPHIIPASPKARTD PPVETTTHHSOGQASQHANSNVNQPGQITSHASRNTPSTAPQASSSPEKFNTQTVPRL ISQTSETAHINQPASGQVTEPKGIFGTYKPRVLTEPAKPANAGVASRQPEATTTVPKL PINPPTARVFIGTASKLSPAVEESHGATPDAHQSKIDREKYAESRPRRTPHLEEGPRE PHVNTPTSAHINVPSSQGQKTVHGRENPGLQTATPSAPQPTASNPRIQYTLPRTDGRL LHDESEVESTPTEEVKRSPKTQDVSHGPEPDDSRWTAPLGPTIEIHRLEHPQILKNIT SLTVPTPRVTPIPPTNIWIPLSHVNIQHEEITRAKNVLMRFIQNVRRKLQASSDALSE ATARTKELYL"

SEQ ID NO 144

CDS

complement (107637..108146)

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	/note="capsid protein; ORF 65; similar to Kaposi's
SEQ ID NO 145	sarcoma-associated herpesvirus ORF 65" /translation="MSSLRVKEPIVQGRLEHDYPNHPLVAEMNNLPQGDMSPAQYAIA KRNYLVFLTAKHHYDMYMQKKNGILRKDHLRGLRGKKDASSSISGVLSGSGSAAPSVA PVASTLGSNSFTTISSGPHSLIGSMGPAPGGGGPGSVASSGIGSTSLSPSDATTLDTR RSSQNKKSK"
SEQ ID NO 146	CDS complement (108152109498)
	/note="unknown; ORF 66; similar to Kaposi's sarcoma-associated herpesvirus ORF 66"
SEQ ID NO 147	/translation="MASGRLPNLAEDEAACHGRGSYPAHRWLDGSRLGLDLAASIRSI GLCPECYVCFVTYGLGAWDGRPPKWACTLISAPSFQTALNEIATGWRPDNPPKNGDVR SRLHDIGRSLLEAYAWVLRCICTGVGCPSDEGLSLTAVPRSAWSRYLVVSFQRACCLV CKTLNCRQRFPLVTCLPQHALDLPVLRKKWNGGGCVSMQLNVPSISRRLGANLNESVP GPSDAGLLASLRELAPTVPCGNPFNALLRSLTFRALLSMSRVVLPIGESTETEISRDL GQKVLAYNVLFPCISLPVWSQVVARSVLEKTVPAPRVVVCLECGYCLNFGRGKFETVN FPPTNVFFSRDQKEKQLSICATTGRVYCSYCGGSHMRVISLFEITCVGDPYLRCVLAN NAAHAIRDANSLVSVVVPCLASPDCATGLLKHLRVAELFYLTSSISSLSCGKCNRS"
SEQ ID NO 148	CDS complement (109524110198)
	<pre>/note="tegument protein; ORF 67; similar to Kaposi's sarcoma-associated herpesvirus ORF 67"</pre>
SEQ ID NO 149	/translation="MSSGKRLVDELCDLVVSYLGPSGISLDLERCQDGAPVYAKGGAV
SEQ ID NO 149	PVCTVRLQHGCVYHLEFVYKFWLHKLERLAYPFAPCFVIINNGLATTLKCFLCKPRDA DAQFGKNLPINSDVYLERNSSVSLGQDDFMKFKARLVFSGDLNVYSSMVICRTYFTEH RQVLQFLVVTPKSAKRLKTLLRTVFALTGHSDGLGALRRTGSVARPSGSELKDIGRGE RAAMTN"
SEQ ID NO 150	CDS 110609111982
	/note="glycoprotein; ORF 68; similar to Kaposi's
SEQ ID NO 151	sarcoma-associated herpesvirus ORF 68" /translation="MFVPWQLETLMRHWPSLRGLVEQSFLPGTPDGAFNSPVLIHTQD SLQPASSCRVCSLLFTLVRTFPPPDSFFEDYGWLCLTCLYAPRSWTATLMVAADLLEL THVYFPQCVKDGPVYTAQSILGIDVQLHFFATRCFRPIDREQILHTSHLNFLQTEFIR GMLEGTIPGSFCFKTSWPRTEKDDQQPTVACCSVGRGSHTNRDNRLPEDLEEAFNSTN AEEKPSLLGVFSATWAESQLLGSDTQQADTHLQPSAFPTPEDADQSQGPCLMHPTLNL KTKNHTASICVLCECLAAHPDAGPVLKDLRRDILENMENNVKLVNRISYILNDPDSLS HVRDEHLRGLIKRCSAQEIHKHFFCDPVCVLNTYSHCPAVLFKCPPPEKYKKLKARLA TGEFLDCNRIFDCETLQTLAVLFKGSQLAKIGKTTSLEIIRELGFQLRRHNIQITHPF QTSNLYI"
SEQ ID NO 152	CDS 112004112897
	/note="unknown; ORF 69; similar to Kaposi's
SEQ ID NO 153	sarcoma-associated herpesvirus ORF 69" /translation="MPKQPRSRLASRAPYAPSVRRPDGPQSTRPASRHGSCKSEIMQW KKLVSDTQFFSALTRRHELGVDFLREMGTPICTSKSVMLPLNLKTIAPGRCVSLSSFG HSSNMGFNCSSCTPTDRSAVSLDANALGEDSARKNSELCSVALTFYHHAEKVVQHKGF YLSLLSHSMEVVRKSFTQPGLLYAHLVLKTFGHDPLPIFTVDADERLALWAVFHTRDL HLGETSLRLIMDNLPNYDITVDCIKQTYIMKFTPSRPDNATVTVPVNSICEAVATLDC TDEFREEIQRGTAIINSQGLL"
CEO TO NO 154	CDS complement (119211119735)
SEQ ID NO 154	/note="FLIP; ORF 71; similar to Kaposi's
	sarcoma-associated herpesvirus ORF 71"
SEQ ID NO 155	/translation="MFPHKRLVDFGRHLEADDREAVLWLFDRPASDDTPEGFANGLCP STGEPGIPLPVLLEAVFLVGRLDLVSTFFLLDVGFIIERLRSSPSYFSPYKHLMLSID RQLSERDVKNLVFLTGDQLGRRRNQSPTFFRWLSQMEKAALVSPSNYMVLSDLLQAVS

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RRDVAKVVAANAPG"

SEQ ID NO 156	CDS complement(119794120558) /note="cyclin D homolog; ORF 72; similar to Kaposi's sarcoma-associated herpesvirus ORF 72"
SEQ ID NO 157	/translation="MASVGPVPTGTIDPVLYQDRALSNLLAHEASFVTSTACYGTIQT EVTVGMRVILGTWMRSVARAHQADASVFPLAVSILDRYLECRSIPRRFQRLGAACLF LAGKIRDLNPFKAAFLCFCAAEDFSVADLLKQEKSILKALRWKLEAVLPTDAIGPTLF KSGFTKEQLFALHSQVVESVHKAIVNPATGGLPPSLVAAACALFSLGAAAPPPARLAE AVGVSAATLAAAAESVATTLREFDEDHILSNARGSS"
SEQ ID NO 158	CDS complement (120866122212)
	/note="latent nuclear antigen; ORF 73; similar to Kaposi's sarcoma-associated herpesvirus ORF 73"
SEQ ID NO 159	/translation="MwgsrQhrsgIvsghgLrsscrghcgrrggtreQagrrgrgrgt AAPAAAPAPAPTTSGPQVRAVAEQGHGSDTETATESRHGSSQGSPSGSGSESVIVLG SPTPSPSGSAPVLASGLSPQNTSGSSPASPASHSPPPSPPSHPGPHSPAPPSSHNPSP NQQPSSFLQPSHHDSPEPPEPPTSLPPPDSPGPPQSPTPTSSPPPQSPPDSPGPPQSP TPQQAPSPNTQQAVSHTDHPTGPSRPGPPFPGHTSHSYTVGGWGPPTRAGGVPCLRLR CTSHNSHEDEAPERQQEQEGEERQQQPARPPRPPRPPRYPIPIPYPSSEEEVPRKYRP
	ORRFYROVLGPRIDPPRPGPWCHGVIFCNSDPYSLYRLARCLQFPGIRASSVRVLPDA
	PGSPVIPAFCITVFCQSRGTAKAVKKARRRWERHHPSAPHFQASIVRMDRGLPIQH"
SEQ ID NO 160	CDS 122866123627 /note="R15; similar to Kaposi's sarcoma-associated herpesvirus K14 and ox-2"
SEQ ID NO 161	/translation="MSGGITLTLLLATLATVRCALQTHYAAVPVHSTASLGCVLTTPH DVLIVTWQKQESPSPVNVATYSSEAGTVVQPPFAGRVDIPEHKLTRTTLKFFNATLED EGCYLCIFNAFGVGKLSGTACLTVYVPLSMSVTFYPPINPTQLVCRAEASPAPSVNWT
	GVPPELCSEPEVFPRPNGTTLVVGRCNVTSVDPEDLENATCLVTHIGGLAAARPLDPV FSDPLEGTSHYVVGVVAAAAVLGIFLTGVFLYRSM"
SEQ ID NO 162	CDS 123924124952
	/note="G protein coupled receptor; ORF 74; similar to Kaposi's sarcoma-associated herpesvirus ORF 74"
SEQ ID NO 163	/translation="MDALNNNLNLMDFLSNYSNSYSSYDDNMSYTLDTESTLCRLTV VFPPTVYAIICFFIFCITLFGNALVLYIFFKFKALANSVDVLMAGLCCNSLFLCASFL FSWLLYVAPQMLTSATCKVEIFFFYLYTYFGVYIVVCISLIRCLLVVFSRRPWVKHGA SGFLCVCVSLIVALALSANASLYRTALRHPETSEWICYEDAGEDTVNWKLRIRTTSAI CGFLVPFGLMVLFYGLTWCMVKSTKLARKGAVRGVIVTVVVLFLIFCLPYHLCNFFDT LLRTGFLAETCYLRDVISVAMHICSLLQSMYSAFVPVVYSGLGSLFRRRVRDTWSVFR CFSTSGSL"
SEQ ID NO 164	CDS complement(125057128953) /note="tegument protein; FGARAT; ORF 75; similar to
	Kaposi's sarcoma-associated herpesvirus ORF 75"
SEQ ID NO 165	/translation="MAQRTNPRWAAAALSPEEEAFIHDNSDAESVLALVPEQCFSEFL LWLVTRPSDNFDNDDDDPALGVIWHLLAPLVNYAPLETRSAHLQGHHTISLPYGPDLM RQPTTRSSEIVQCLRDSGLDRTLRLEVGRHLSCQTRRFVADRVPPGTLAALTLGTLVE YDVRVQRQLPVTLQSTAWRPLPERDPICAAVMLPLQRNILPLAVQASNGNSYTVSRYA VMARRSYSCVFQRLPCENVTHIADSFTHLHSAIQTGAGALQNILFHATLLPGGEIRSA LCGFYATTPSVGAFSRARHRAINTTATLHCQQLARTGTPVLGGFLKTVHSATTSEANV
·	LCGFYATTPSVGAFSRARHRAINTTATHHCQQLARIGIPVLGGFLKIVNSATISLANV ITTTSLLSCVPQAYTFLRRSLFSQPIICLGSFEPVDGDGNQRSLYLGSAAGITRITQT LSLAYEILEGPLFTSINRAHEPASVIGHLGALVSRGGLRLFVSQLPPTILSQLTATPD ISRETVNDILVNKFLNVSACVVFAVLPRDTEPEPGPLDAIRRAARICGCPFAVVGETC EELGIQFVNDLELWNPGAWPIRQPTSAEVIATFGFDEQPVSSNWLVRPEEPEDGGEQA

PSPTDWGLFRLASVVDQLLRCPTVGSKEFVTRHVDRCSNGLVAQQCEVGPLGRPLSDY

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HIVNHTSVFTDRMARVPIYRPQPITRQDATERLVSPETWVTQGRGRNRWVGQCVAYGE QAYKMGINAAVGARYAICEAVTNIMLAHVRRLSDITLTASVGWNPEDDQAWLLQHTLF ACKELCRDLSINFAITSAGSTPCLSEELISATQQHQTVAPVPFNAVVITATAEVKSSR QRVTPDLKATGNLIVLVSFPVPHLTQGSTFEHLCLLPSPTLPDVQATHLANLFMLTEA LLSRGLVVSGHDVSDGGMVVTAIEMALAGNRGLQIRIPSEETPLQWLVSETPGVIFEI QPQHVDEVRQACQNFDCRATVCGTVGQEGLSERIVISHNNDEVYSQTLTSVAANWTSF SDEQWYSWGPSFTPAQELYRKDYGCNQHNLGHLAEVCRNSELTLFATPSRPPAVAALV TPGAPLPRALMAAFTNVGFDVAAVSTDDLRGGNILRGFSGLTIGGTVGIEDSYVGARC AIMGLLNDPGCYGGLMAFFRRADTFSLCCGEFGFQLLGALGLLRETPHDTPGPKTPDQ WDIHLEENASGNHECLWLNLHIPQTTISIMFRVLRGLVLPGWANGRYLGVRYPRDAME YHLNQQQRIALNYHTGNADPRMFAQHYPRNPSANSAVAAITSPDGRHLASLVDPAVTF HPWQWAYVPPELADMTVSPWALAFQSLFLWCIRNRQ"

SEQ ID NO 166 PCR primer CCTATGGGCTCCATGAGC

SEQ ID NO 167 PCR primer ATCGTCAATCAGGCTGCG

SEQ ID NO 168 PCR primer ATATTAAACACTCGCCGC

SEQ ID NO 169 PCR primer ATGAGGGGCCTTTTCGTGTGC

SEQ ID NO 170
PCR primer
CTGAATCCCGCTGCCAAGGCC

SEQ ID NO 171
PCR primer
ATGTTCCCTGTCTGGTTCGTC

SEQ ID NO 172 PCR primer TTACATCATAGCTATTGCGCG

SEQ ID NO 173 <213> Macaca mulatta rhadinovirus 17577

nucleotides complement(23398..23668)

SEQ ID NO 174 <213> Macaca mulatta rhadinovirus 17577

nucleotides 25065..25368

SEQ ID NO 175 <213> Macaca mulatta rhadinovirus 17577

nucleotides 25518..26525

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SEQ ID NO 176 <213> Macaca mulatta rhadinovirus 17577

nucleotides 114979..115383

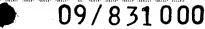
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SEQ ID NO 178
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SEQ ID NO 179
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nucleotides 132333..133719
partial terminal repeat





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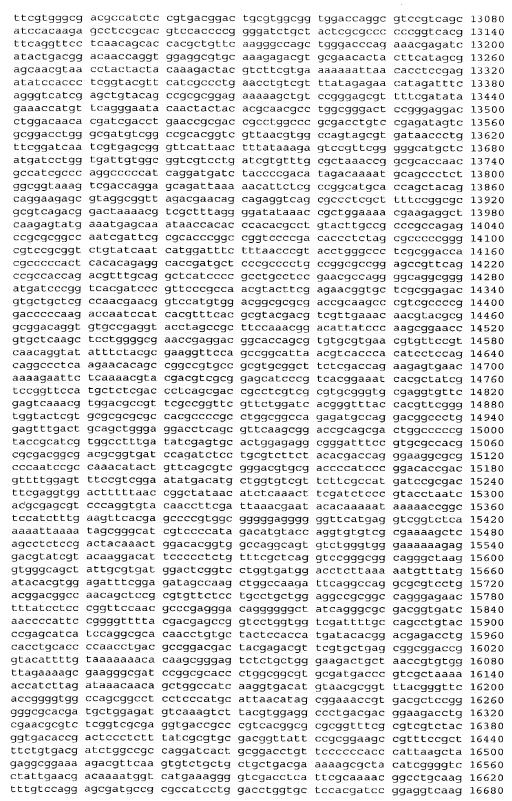
SEQUENCE LISTING

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<110> Oregon Health Sciences University
<120> Cloning of Rhadinovirus Genome and Methods for its Use
<130> 53683
<140> PCT/US99/26260
<141> 1999-11-05
<150> 60/107,507
<151> 1998-11-06
<150> 60/109,409
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Val His Cys Asp Glu Asn Cys Lys Pro Pro His Phe Thr Glu Tyr Arg

20 25 30

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Thr Glu Cys Leu Gln Asn Gly Thr Trp Ser Thr Pro Asn Phe Pro Cys
65 70 75 80

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Ser Thr Pro Met Pro Glu Thr Pro Met Pro Glu Thr Pro Thr Pro Asp Tyr Gln Lys Ile Asn Leu Ser Thr Ala Lys Thr Ala Thr Thr Pro Asn Ala Phe Val Thr Thr Val Val Ser Pro Glu Lys Asp Asp Val Thr Cys Val Lys Pro His Phe Glu Arg Phe Met Val Lys Ala Glu Asn Asp Lys Glu Lys Tyr Ser Val Gly Ala Ser Val Glu Leu Ile Cys Arg Pro Gly Phe Thr Lys Met Gln Ser Thr Val Ser Val Glu Cys Leu Ser Asn Gly Thr Trp Thr Ala Pro Asn Ala Lys Cys His Arg Lys Lys Cys Pro Thr Pro Gln Glu Leu Leu Asn Gly Glu Tyr Ile Val Thr Ser Gly Glu Asp Ala Phe Lys Tyr Gly Thr Asn Ile Thr Tyr Lys Cys Asn Glu Gly Tyr Gln Leu Leu Gly Ser Met Val Arg Ile Cys Met Leu Lys Asp Asp Leu Lys Thr Val Asp Trp Glu Pro Lys Ala Pro Ile Cys Asp Ile Glu Lys Cys Lys Pro Pro Pro Gln Ile Thr Asn Gly Lys Tyr His Pro Val Lys 450 455 Asp Phe Tyr Gln Tyr Leu Asp Thr Val Thr Phe Ser Cys Asn Arg Asp Phe Ser Leu Val Gly Asp Glu Met Thr Thr Cys Ile Ser Asn Thr Trp Asn Lys Pro Phe Pro Arg Cys Glu Gln Ile Thr Cys Ser Ala Pro Asn Ile Ala His Gly Lys Leu Leu Thr Gly Ser Ser Ser Val Tyr Lys Tyr Gly Gln Ser Val Thr Ile Gly Cys Glu Thr Gly Phe Thr Leu Ile Gly Ser Glu Ile Ser Thr Cys Lys Asp Ser Ser Trp Asp Pro Pro Leu Pro Thr Cys Val Pro Ala Val Ser Met Pro Ser Asp Thr Pro Lys Pro Glu Thr Lys Lys Pro Asn Thr Pro Thr Pro Glu Ala Pro Lys Pro Asn Thr Pro Asn Val Gly Thr His Thr Pro Phe Lys Pro Pro Pro Gln Asn Pro Pro Ile Ala Pro Pro Met Ser Lys Trp Lys Arg His Val Val Leu Val Leu Phe Ala Ser Val Ala Ser Leu Leu Phe Val Leu Ala Ala Leu Tyr Cys Cys Phe Leu Lys

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_	tac Tyr	_		_	_		_				_	_			624
	tac Tyr 210			_	_	_				_					672
	atc Ile	-	_	_	_	_			_	 _		_			720

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gtt Val	ccg Pro	aaa Lys	tca Ser	att Ile	aaa Lys	ata Ile	aaa Lys	aat Asn	aga Arg	atc Ile	att Ile	ttt Phe	tcc Ser	aac Asn	acc Thr	2208

osgados osogol

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465 Pro	Ala	Val	Thr	Thr	470 Pro	Gln	Arg	Arg	Asp	475 Pro	Tyr	Val	Val	Thr	480 Gly
Thr	Ala	Gly	Thr	485 Phe	Asn	Asp	Leu	Glu	490 Ile	Leu	Gly	Asn	Phe	495 Ala	Ser
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Gly	Glu	Val	Met 580	565 Lys	Phe	Val	Asn	Ser 585	570 Met	Ile	Lys	Asn	Asn 590	575 Phe	Asn

Phe Arg Glu His Val Lys Ser Val His His Ile Leu Gln Phe Cys Cys Asn Val Tyr Trp Gln Ala Pro Cys Ala Val Phe Leu Asn Leu Tyr Tyr Lys Ser Leu Leu Trp Ile Ile Gln Asp Ile Cys Leu Pro Tyr Cys Met Ile Tyr Glu Gln Asp Asn Pro Ala Met Gly Ile Leu Pro Ser Glu Trp Leu Lys Met His Phe Gln Thr Leu Trp Thr Asn Phe Lys Ala Ala Cys Leu Asp Arg Gly Val Leu Thr Gly Cys Glu Leu Lys Ile Val His Arg Asp Met Phe Cys Asp Phe Phe Asp Thr Asp Ala Gly Ser Asn Gly Leu Met Ala Pro Phe Lys Met Gln Val Arg Ile Ala Arg Ala Met Met Val Val Pro Lys Ser Ile Lys Ile Lys Asn Arg Ile Ile Phe Ser Asn Thr Ala Gly Ser Glu Ala Val Gln Ser Gly Phe Val Lys Pro Thr Gly Thr Arg Asp Thr Tyr Val Val Ala Gly Pro Tyr Met Lys Phe Leu Asn Ser Leu His Arg Ala Leu Phe Pro Asp Thr Lys Thr Ala Ala Leu Tyr Leu Trp His Lys Ile Ser Gln Thr Asn Lys Thr Pro Val Leu Lys Asp Val Pro Asp Asp Glu Leu Ala Glu Leu Val Ser Tyr Val Lys Thr Asn Ser Leu Ala Phe Glu Glu Thr Asn Val Leu Asp Val Val Pro Asp Ser Leu Met Ser Tyr Ala Arg Ile Lys Leu Asn Gly Ala Ile Leu Arg Ala Cys Gly Gln Ile Gln Phe Tyr Ala Thr Thr Leu His Cys Leu Thr Pro Val Leu Gln Thr Ile Asp Ala Glu Glu Tyr Pro His Val Leu Gly Ser Ala Ala Ile Ala Thr Pro Val Ala Tyr Leu Ala Glu Ile Arg Gly Arg Thr Ala Leu Thr Val Gln Thr Thr Ala Arg Gln Pro Val Ala Ala Thr Gly Arg Leu Arg Pro Val Ile Thr Val Pro Met Val Val Asn Lys Tyr Thr Gly Val Asn Gly Asn Asn Val Phe His Cys Gly Asn Leu Gly Tyr Phe Ala Gly Arg Gly Val Asp Arg Asn Leu Trp Pro Glu Ser Ser Pro Phe Lys Lys Thr Gly Val Ser Ala Met Leu Arg Lys Arg His Val Met Met Thr Pro Ile Ile Asp Arg Leu Ile Lys Arg Ala Ala Gly Gln Thr Ile Ser Thr Phe Glu Ala Glu Ser Val Lys Arg Ser Val Gln Ala Leu Leu Glu Asp Lys Asp Asn Pro Asn Leu Leu Lys Ser Val Ile Leu Glu Leu Ile Arg His Leu Gly Lys Gly Cys Gln Asp Leu Ser Ser Glu Asp Val Gln Tyr Tyr Leu Gly Asp Tyr Cys Met Leu Thr Asp Glu Val Leu Phe Thr Leu Asp Asn Ile Ala Gln Ser Gly Val Pro Trp Thr Ile Glu Asp Ala Gly Ala Leu Ile Glu Asp Arg Gln Asp Ala Asp Asp Leu Gln 1075 1080 1085

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cca Pro	aat Asn	cag Gln 195	gly ggg	tcc Ser	agt Ser	tta Leu	ctc Leu 200	gcg Ala	gtg Val	ttg Leu	gca Ala	gac Asp 205	cga Arg	cac His	tgc Cys	624
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tcc cta gc Ser Leu Al 625	g cgc gag a Arg Glu	aaa ctc Lys Leu 630	tcc atc Ser Ile	tcc aac Ser Asn 635	cta gac Leu Asp	gtt aaa Val Lys	ggc 1920 Gly 640
ctg acg tc Leu Thr Se	c ggc ctg r Gly Leu 645	tat cta Tyr Leu	acg tac Thr Tyr	gag caa Glu Gln 650	gac gcg Asp Ala	ccg ctc Pro Leu 655	Val

cta att tct caa aat acc ggc tgg ata ttt aaa gac ctg tac gct ctt Leu Ile Ser Gln Asn Thr Gly Trp Ile Phe Lys Asp Leu Tyr Ala Leu ctg tac cat cac ctg caa ctg tcc gac ggc cat gat gat aac taa Leu Tyr His His Leu Gln Leu Ser Asp Gly His Asp Asp Asn <210> 11 <211> 686 <212> PRT <213> Macaca mulatta rhadinovirus 17577 <400> 11 Met Ala Arg Glu Leu Ala Ala Leu Tyr Ala Gln Leu Ser Ala Leu Ala Val Asp Leu Ser Leu Val Ile Phe Ala Asp Pro Arg Ser Ile Asp Gly Ala Arg Ile Leu Lys Thr Lys Thr Gln Ile Glu Asn Leu Asn Arg Asp Leu Leu Pro Leu Leu Arg Glu Gln Asn Ser Val Glu Thr Ser Ser Leu Ser Leu Glu Val Glu His Leu Ala Lys Asn Ile Glu Asp Lys Leu Gly Glu Leu Glu Arg Ser Leu Arg Gln Arg Tyr Ser Ser Arg Glu His Phe Glu Thr Leu His Leu Arg Pro Glu Cys His Tyr His Ser Thr Val Thr Phe Gln Phe Tyr Gly Gly Gly Leu Ile Asp Val Asn Met Cys Leu Ile Asn Asp Val Glu Leu Leu Cys Lys Arg Leu Gly Ser Val Phe Tyr Cys Ile Gly Ala Asn Glu Ala Leu Ser Gly Leu Asn Arg Val Leu Thr Phe Leu Ser Thr Leu Arg Gly Ile Ser Pro Ile Pro His Pro Asp Leu Tyr Val Thr Ser Val Pro Cys Val Gln Cys Leu Arg Glu Ile Glu Leu Val Pro Asn Gln Gly Ser Ser Leu Leu Ala Val Leu Ala Asp Arg His Cys Asp His Leu Cys Lys Lys Val Arg Ala Glu Pro Ile His Gly Leu Phe Glu Thr Glu Leu Ser Gln Leu Gly Leu Lys Val Thr Lys Arg Ser Asp Ala Thr Gln His Gly Val Arg Ser Ser Ala Asp Gln Leu Arg Glu Ser Ser Leu Ala Ala Ile Gln Asp His Asn Ile Phe Lys Arg Val Ser Ala Ser Ile Met Glu Leu Ser Asn Leu Ile Tyr Trp Asn Ala Gly Gln Thr Gly Leu Gln Thr Gly Thr Glu Asn Glu Cys Ser Gln Met Ala Arg Leu Leu Thr His Glu Ala Asp Met His Glu His Arg Ala Leu Ile Thr Pro Lys Leu Ser Ala Thr His Phe Tyr Asp Cys Phe Arg Pro Asp Pro Ile

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Tyr Thr Asn Val Met Arg Lys Gln Asn Glu Leu Phe Thr Arg Leu Asn
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		aaa Lys			_	_		_	_	_		-	336
		gcc Ala 115											384
		gg¢ Gly											432
		atg Met											480
_		ggc Gly											528
		ttt Phe	_		-	_	 					_	576
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_		_								gcg Ala		_	_	_		1344
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_	_	_	_					_	_	atg Met	_					1488
	_				_				_	gcc Ala				_	_	1536
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tac Tyr	ccc Pro	gac Asp	ata Ile	gac Asp	aaa Lys	atg Met	cag Gln	ccc Pro	tct Ser	ggc Gly	ggt Gly	aaa Lys	gtc Val	gac Asp	cag Gln	2304

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288

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<213> Macaca mulatta rhadinovirus 17577

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ggg gtg gac Gly Val Asp	cag ctg cgt Gln Leu Arg 180	tac gtg gtg Tyr Val Val 185	gat cta att Asp Leu Ile	aac agg cgg Asn Arg Arg 190	ccc 576 Pro
			tgg aac ccc Trp Asn Pro		
cgg atg gct Arg Met Ala 210	ctc cct cct Leu Pro Pro	tgt cac gtt Cys His Val 215	ttg tgt cag Leu Cys Gln 220	ttt tac gtg Phe Tyr Val	gct 672 Ala
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			cgg ctg aag Arg Leu Lys 300		
gtg gcg cgt Val Ala Arg 305	ctg gag gac Leu Glu Asp 310	ttt acg cgc Phe Thr Arg	gcg gat ctg Ala Asp Leu 315	agt ctc gag Ser Leu Glu	ggc 960 Gly 320
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35 Gly Glu Leu 50	Gln Tyr Leu	40 Ala His Leu 55	Asp Leu Ile 60	45 Ile Lys His	Gly

Val Gln Arg Glu Asp Arg Thr Gly Val Gly Thr Arg Ser Val Phe Gly 70 75 Leu Gln Ala Arg Tyr Asn Leu Arg Asp Glu Phe Pro Leu Leu Thr Thr 95 90 8.5 Lys Arg Val Phe Trp Arg Gly Val Val Glu Glu Leu Leu Trp Phe Ile 105 100 Arg Gly Ser Thr Asp Ser Thr Glu Leu Ser Arg Arg Gly Val Lys Ile 120 125 Trp Asp Ala His Gly Ser Arg Ala Phe Leu Ala Ala Gln Gly Phe Gly 140 135 Asp Arg Arg Glu Gly Asp Leu Gly Pro Val Tyr Gly Phe Gln Trp Arg 150 155 His Phe Gly Ala Glu Tyr Arg Gly Ala Asp Ala Asn Tyr Glu Gly Gln 175 165 170 Gly Val Asp Gln Leu Arg Tyr Val Val Asp Leu Ile Asn Arg Arg Pro 180 185 190 His Asp Arg Arg Ile Val Met Cys Ala Trp Asn Pro Ala Asp Leu Ala 200 195 Arg Met Ala Leu Pro Pro Cys His Val Leu Cys Gln Phe Tyr Val Ala 220 215 Arg Gly Glu Leu Ser Cys Gln Leu Tyr Gln Arg Ser Ala Asp Met Gly 235 Leu Gly Val Pro Phe Asn Ile Ala Ser Tyr Ala Leu Leu Thr Tyr Leu 255 245 250 Ile Ala His Val Thr Gly Leu Thr Pro Gly Asp Phe Val His Thr Leu 260 265 Gly Asp Ala His Val Tyr Asn Asn His Val Asp Pro Leu Leu Gln 275 280 285 Leu Arg Arg Thr Pro Arg Pro Phe Pro Arg Leu Lys Ile Leu Arg Lys 295 300 Val Ala Arg Leu Glu Asp Phe Thr Arg Ala Asp Leu Ser Leu Glu Gly 310 315 Tyr Asp Pro His Pro His Ile Glu Met Glu Met Ala Val 330 325

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Val Asp Tyr Ala Phe Pro Met Gly Ser Met Ser Gly Pro Ala Pro Glu
ctc tgc tgt ttg ggg tat gta act cat ctg ccg cca ccc ggt tta gtg
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Leu Cys Cys Leu Gly Tyr Val Thr His Leu Pro Pro Pro Gly Leu Val
qte tet tae tee cae ace teg teg cag tge teg gtg gae gee gtg ata
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Val Ser Tyr Ser His Thr Ser Ser Gln Cys Ser Val Asp Ala Val Ile

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Asn Ser Leu Pro Val Asp Val Tyr Ala Ile Glu Gly Ile Phe Leu Tyr 25

20

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Ile	Lys 50		Phe	Ile	Ser	Ala 55		Leu	Lys	Asp	Ser 60	Ala	Arg	Leu	Tyr	
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	Ile	Gly	Gln	Gly 85	Met	Leu	Gln	Ile	Asn 90	Thr	Asp	Gly	Arg	His 95	Asn	
Trp	Gly	Arg	Ala	Leu	Ala	Val	Leu	Gly	Leu	Gly	Ala	Tyr	Val	Val	Asp	

Lys Val Lys Asp Asp Glu Arg Leu Leu Thr Phe Ala Ile Ala Val Leu 120 125 115 Pro Val Tyr Ala Tyr Glu Ala Leu Glu Ser Gln Trp Phe Arg Ser His 135 140 Gly Glu Trp Glu Gly Leu Arg Asn Tyr Cys Glu Arg Ile Leu Arg His 150 155 145 Arg Arg Asn Ala Arg Arg His Met Cys Tyr Gly Val Ala Ala Gly Leu 170 165 Leu Ala Leu Val Ala Leu Phe Ala Ile Arg Arg 180 <210> 28 <211> 1611 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(1611) <400> 28 48 atg act ecc gtg tac gtt ggg gga tac gtg gac gtg gtc agc cta cca Met Thr Pro Val Tyr Val Gly Gly Tyr Val Asp Val Val Ser Leu Pro aag ata gaa aag gag ctg tat tta gag ccc tca atc gtg gcg acc ctg Lys Ile Glu Lys Glu Leu Tyr Leu Glu Pro Ser Ile Val Ala Thr Leu 2.5 2.0 144 ctc ccg tat acg gac cct cta ccg ata aac ata gag cac gtc ccc gaa Leu Pro Tyr Thr Asp Pro Leu Pro Ile Asn Ile Glu His Val Pro Glu 40 qcc cac gta ggt cac aca atc ggt ctc ttc caa gta aca cac ggg ata 192 Ala His Val Gly His Thr Ile Gly Leu Phe Gln Val Thr His Gly Ile 55 240 ttt tgc ctg ggc aag cta acg agc cac gat ttc ttg gcc ctg gcg tca Phe Cys Leu Gly Lys Leu Thr Ser His Asp Phe Leu Ala Leu Ala Ser 75 egg etc geg ggg gac teg ega geg gee eag atc eag eta aac eec atg 288 Arg Leu Ala Gly Asp Ser Arg Ala Ala Gln Ile Gln Leu Asn Pro Met 8.5 90

105

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_			_	_	gag Glu	_	_				_	_				528
_			_	_	cag Gln					_						576
			_		aag Lys			-					-	-	_	624
	_	_			aca Thr	_				_	_	_		-	_	672
			_	_	agc Ser 230	_			_	_	_					720
_	_		_		atg Met	_	_		_		_					768
	_	_			ctg Leu		_		_	_	_		_		_	816
					cgc Arg											864
_	_				cca Pro	_			_	_			_		-	912
			_	_	ccc Pro 310	_			_				_	_		960
	_				tct Ser	_			_	_				_	_	1008
				_	gca Ala	_		_		_	_	_			_	1056
	_				gag Glu											1104
	_	_			gcg Ala						-			-		1152
_	-				aaa Lys	_	_			_				_	_	1200

385		390				395					400	
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cag acc t Gln Thr T	ac gcg yr Ala 420	tct gcg Ser Ala	ccg ta Pro Ty	ac cta yr Leu 425	gcg Ala	tac Tyr	cag Gln	ccg Pro	cag Gln 430	tgg Trp	tat Tyr	1296
tcc gga a Ser Gly T	cg gac hr Asp 35	acc cat Thr His	ctc ca Leu H:	is Ala	cca Pro	cag Gln	ccg Pro	tac Tyr 445	cag Gln	agc Ser	gcg Ala	1344
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cac gcc g His Ala G 465	gt ctc ly Leu	gcc acg Ala Thr 470	cag co Gln Pi	eg gca ro Ala	act Thr	cca Pro 475	gcc Ala	ccc Pro	gcc Ala	gcc Ala	caa Gln 480	1440
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ggc gcg to Gly Ala C	gc ccg ys Pro 500	ccc ctc Pro Leu	gat co	ca gaa co Glu 505	tgc Cys	gga Gly	cag Gln	tcc Ser	gcg Ala 510	cgg Arg	gcc Ala	1536
ccg gtg g Pro Val G 5	ag gcc lu Ala 15	agc gca Ser Ala	cag co Gln Pr 52	co Ala	ccc Pro	gtg Val	tca Ser	cag Gln 525	ata Ile	caa Gln	aaa Lys	1584
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Leu Pro T		Asp Pro		o Ile	Asn	Ile	Glu	His 45	Val	Pro	Glu	
Ala His V		His Thr	Ile Gl	ly Leu	Phe	Gln	Val 60	Thr	His	Gly	Ile	
Phe Cys Lo	eu Gly	Lys Leu 70	Thr Se	er His	Asp	Phe 75	Leu	Ala	Leu	Ala	Ser 80	
Arg Leu A	la Gly	Asp Ser 85	Arg Al	a Ala	Gln 90	Ile	Gln	Leu	Asn	Pro 95	Met	
Pro Arg A	sp Pro 100		Glu Me	et Leu 105	His	Thr	Trp	Leu	Pro 110	Glu	Leu	
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Pro Ser Tyr Ala Pro Pro Val Ala Pro Pro Phe Pro Phe Gln Ser Ala
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His Lys Asp Val Met Ala Leu Ser Lys Asn Ile Leu Asp Ile Gln Ala
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Asp Leu Arg Asp Leu Lys Arg Ala Ala Ser Gln Thr Ser Gly Ala Gln
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Asp Ala Asp Gln Arg Pro Gln Pro Pro Pro Val Gln Phe Ser Trp Pro
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Gln Gly Ile Gln Gln Thr Gln Pro Pro Pro Gln Pro Ala Ser His
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His Ala Gly Leu Ala Thr Gln Pro Ala Thr Pro Ala Pro Ala Ala Gln
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<212> DNA

<213> Macaca mulatta rhadinovirus 17577

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														acg Thr		672
														ggt Gly		720
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														ctg Leu		816
														gcg Ala		864
														cac His		912
														aag Lys		960
														ctt Leu 335		1008
														caa Gln		1056
		-	_											gtt Val		1104
		-		_	_									cgg Arg		1152
														gaa Glu		1200
_									_					gat Asp 415		1248
		_								-				gac Asp		1296

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150

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Leu Asn Lys Glu Thr Lys Tyr Asn Ala Asp Phe Phe Thr Asn His Val
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Pro Gly Thr Gly Gln Pro Pro Gly Pro Asp Leu Val Tyr Ile Leu Ala
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Thr Thr Leu Phe Ser Glu Asp Val Pro Pro Phe Gln Ala Tyr Gln Trp
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Leu Thr Glu Asn Tyr Ile Ser Pro Ile Leu Ser Arg Ala Pro Asp Ala
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Lys Ala Ile Ser His Leu Lys His Thr Arg Pro Phe Val Asn Leu Thr
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465 470
Phe Arg Asp Ala Gly Ser Leu Ile Gln Ala Gln Thr Ser Leu Arg Leu
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Thr Ala Glu Glu Gly Leu Ala Ala Ile Leu Ser His Pro Ser Pro Pro
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<211> 1053

<212> DNA

<213> Macaca mulatta rhadinovirus 17577

<220>

<221> CDS

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tca a						-	-				_				_	192
ttt t Phe I 65			_					-		-	-	_			_	240
gtt a Val A			_				_	_		_		_		-		288
tgc (Cys I					_					_						336
cga t Arg T		_					-		_		_	_				384
aga g Arg <i>l</i>	_			-			_			_	_					432
cta t Leu I 145		_		-												480
gcg t Ala I		-								_						528
ttt t Phe I		_			_	_			_	_		_				576
ata d Ile I		_			_		_			_			_			624
cca t Pro S				_									_	_	_	672
aag a Lys I 225																720

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144

192

240

288

200 195 Pro Ser Lys Gln Arg Ser Val Tyr Ser Gln Thr Ile Ser Asp Arg Arg 220 210 215 Lys Lys Lys Arg Val Cys Asp Ala Lys Ser Thr Ala Gly Ala Lys Gly 230 235 Ser His Ala Ala Lys Lys Pro Ala Pro Ala Arg Thr Arg Gln Arg Ala 250 245 Ala Asn Ala Pro Thr Gly Asn Arg Ser Gly His Ala Arg Pro Arg Asn 265 260 Asn Ser Lys His Gly Arg Gly Ser Ala Val Pro Gly Gln Gly Asn Arg 280 285 Gln Cys Pro Asn Ile Thr Lys Pro Ala Thr Gln Asn Arg Pro Ala Asp 295 Thr Trp Arg Arg Val Arg Cys His Asn Ser Pro Arg Arg Pro Gly Ile . 315 310 His Gly Lys Pro Gly Ser Pro Ser Gly Ala Pro Ala Lys Pro Val His 325 330 Glu Pro Lys Pro Met Ala Ala Thr Ile Arg Ala Val Val Gln

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cgt Arg 145	act Thr	act Thr	aca Thr	cgc Arg	ccg Pro 150	gtt Val	gaa Glu	agc Ser	ggt Gly	gac Asp 155	aag Lys	aga Arg	aat Asn	ttc Phe	acc Thr 160	480
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att Ile	caa Gln 370	tac Tyr	gca Ala	cca Pro	aaa Lys	gat Asp 375	aga Arg	gat Asp	tac Tyr	aat Asn	ttt Phe 380	att Ile	ttt Phe	aac Asn	gcc Ala	1152
							gaa Glu									1200
							gat Asp									1248
							aat Asn									1296
							ggt Gly 440									1344
							gta Val									1392
							ttc Phe									1440
gct Ala	gca Ala	att Ile	aac Asn	gtt Val 485	aac Asn	ata Ile	agc Ser	gly ggg	gac Asp 490	atg Met	ctg Leu	cac His	ttt Phe	atg Met 495	ttc Phe	1488
gct Ala	atg Met	gga Gly	aac Asn 500	ctc Leu	agg Arg	tgc Cys	ttt Phe	tta Leu 505	ccg Pro	gtg Val	aag Lys	cac His	att Ile 510	ttc Phe	ccg Pro	1536
gtt Val	tcg Ser	att Ile	gcg Ala	aac Asn	tgg Trp	aac Asn	tct Ser	acg Thr	tta Leu	gac Asp	ctc Leu	cac His	ggg Gly	ctt Leu	gaa Glu	1584

9,

į.	515		520				525				
aac caa t Asn Gln 1 530	tac ata q Tyr Ile Y	Val Arg A	cgg ggg Arg Gly 535	cgg c	ega g Arg A	ac gtt sp Val 540	ttc Phe	tgg Trp	acc Thr	act Thr	1632
aat ttc o Asn Phe 1 545					Asp G						1680
ttt aag g Phe Lys A	Ala Ala '			Ser I							1728
ctt aaa a Leu Lys l											1776
gct aga a Ala Arg :											1824
aga aat a Arg Asn 1 610	Lys Ala (Gln Ile (Gln Thr 615	Leu F	His I	ys Arg 620	Phe	Ile	Glu	Cys	1872
ttg gtg g Leu Val G					Leu A						1920
cgc gcc g	Ala Arg I	Leu Gly ' 645	Thr Phe	Asp I	Phe S 650	Ser Lys	Arg	Ile	11e 655	Ser	1968
cac acc a											2016
Asn Leu	Ile Pro : 675		Tyr Val 680	Arg :	Ser I	Lys Lys	1le 685	Arg	Leu	Asp	2064
gag tta g Glu Leu 690	gga cgc (Gly Arg)	Asn Ala	aat ttt Asn Phe 695	atg Met S	tct t Ser I	ttc ata Phe Ile 700	: Ala	acg Thr	acc Thr	ggt Gly	2112
cac gcg His Ala 705	Phe Ser A	Asn Leu 1 710	Lys Pro	Gln '	Val 1	Ile Arq 715	y His	Thr			2160
cgt ttg	Gly Leu 1	cac tgg His Trp 725		Lys 2							2199

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<211> 732

<212> PRT

<213> Macaca mulatta rhadinovirus 17577

<400> 43

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490 Ala Met Gly Asn Leu Arg Cys Phe Leu Pro Val Lys His Ile Phe Pro 510 505 500 Val Ser Ile Ala Asn Trp Asn Ser Thr Leu Asp Leu His Gly Leu Glu 520 Asn Gln Tyr Ile Val Arg Arg Gly Arg Arg Asp Val Phe Trp Thr Thr 535 540 Asn Phe Pro Ser Val Val Ser Ser Lys Asp Gly Cys Asn Val Ser Trp 555 550 Phe Lys Ala Ala Thr Ala Thr Ile Ser Lys Ile Tyr Gly Arg Pro Leu 565 570 Leu Lys Lys Leu Ser Asp Glu Leu Asn Pro Ile Leu Ser Val Pro Tyr 585 Ala Arg Ile Asp Gln Val Lys Asn Thr Ile Phe Thr Thr Leu Glu Thr 600 605 Arg Asn Lys Ala Gln Ile Gln Thr Leu His Lys Arg Phe Ile Glu Cys 620 615 Leu Val Glu Cys Cys Ser Phe Leu Arg Leu Asp Leu Gly Ala Leu Asn 630 635 Arg Ala Ala Arg Leu Gly Thr Phe Asp Phe Ser Lys Arg Ile Ile Ser 650 645 His Thr Lys Ser Lys His Glu Cys Ala Ile Leu Gly Tyr Lys Lys Cys 665 670 660 Asn Leu Ile Pro Lys Ile Tyr Val Arg Ser Lys Lys Ile Arg Leu Asp 685 680 Glu Leu Gly Arg Asn Ala Asn Phe Met Ser Phe Ile Ala Thr Thr Gly 695 700 His Ala Phe Ser Asn Leu Lys Pro Gln Val Ile Arg His Thr Ile Arg 715 710 Arg Leu Gly Leu His Trp Arg His Lys Ala Lys Ile 725

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Lys 65	Phe	Leu	Glu	Thr	Ser 70	Leu	Ala	Val	Ala	Cys 75	Val	Asn	Thr	Glu	Phe 80	
aag Lys	gac Asp	ctc Leu	aaa Lys	cga Arg 85	atg Met	acg Thr	gat Asp	gga Gly	aaa Lys 90	att Ile	cag Gln	ttt Phe	aag Lys	gta Val 95	tct Ser	288
gta Val	ccg Pro	acc Thr	atc Ile 100	gcg Ala	tat Tyr	ggg ggg	gac Asp	ggc Gly 105	agg Arg	cgg Arg	ccc Pro	aca Thr	aaa Lys 110	caa Gln	aaa Lys	336
caa Gln	tac Tyr	att Ile 115	atc Ile	atg Met	aag Lys	gcc Ala	tgc Cys 120	aat Asn	aag Lys	cat His	cac His	att Ile 125	ggt Gly	gcc Ala	gag Glu	384
ata Ile	gag Glu 130	ctg Leu	tcg Ser	act Thr	gat Asp	gac Asp 135	atc Ile	gag Glu	ctg Leu	cta Leu	ttc Phe 140	att Ile	gac Asp	aga Arg	gaa Glu	432
acc Thr 145	Pro	ctc Leu	gat Asp	tac Tyr	aca Thr 150	gaa Glu	tac Tyr	gcc Ala	Gly ggg	gcc Ala 155	gta Val	aag Lys	acg Thr	att Ile	acc Thr 160	480
gcc Ala	tct Ser	ctc Leu	cag Gln	ttt Phe 165	ggc Gly	gtg Val	gac Asp	gcg Ala	ctg Leu 170	gag Glu	agg Arg	ggc Gly	ctg Leu	gta Val 175	gat Asp	528
acc Thr	gta Val	ttg Leu	aat Asn 180	gtt Val	aag Lys	ctt Leu	agg Arg	tçc Ser 185	gcc Ala	ccg Pro	ccg Pro	atg Met	ttt Phe 190	att Ile	cta Leu	576
aaa Lys	aca Thr	cta Leu 195	tca Ser	gac Asp	ccg Pro	gtc Val	tac Tyr 200	acc Thr	gaa Glu	cgg Arg	ggt Gly	cta Leu 205	aag Lys	aag Lys	gct Ala	624
gtt Val	aag Lys 210	tca Ser	gac Asp	atg Met	gtg Val	tcc Ser 215	atg Met	ttc Phe	aaa Lys	agc Ser	tac Tyr 220	ctt Leu	atg Met	gat Asp	aac Asn `	672
tcg Ser 225	ttt Phe	ttc Phe	ctc Leu	gac Asp	aaa Lys 230	tca Ser	gac Asp	atc Ile	gcc Ala	gtc Val 235	aag Lys	gga Gly	aag Lys	cag Gln	tac Tyr 240	720
gtg Val	ctg Leu	tcg Ser	gtt Val	ctc Leu 245	tcc Ser	gac Asp	atg Met	gtg Val	ggg Gly 250	gcg Ala	gtg Val	tgt Cys	cac His	gaa Glu 255	acg Thr	768
gtt Val	ttt Phe	aag Lys	999 Gly 260	acg Thr	aat Asn	acg Thr	tat Tyr	ctg Leu 265	Ser	gca Ala	tcg Ser	gga Gly	gag Glu 270	cca Pro	att Ile	816
gcc Ala	gga Gly	gtc Val 275	atg Met	gag Glu	acc Thr	acg Thr	gaa Glu 280	Asn	gta Val	atg Met	cga Arg	aaa Lys 285	Leu	tta Leu	aac Asn	864
atg Met	cta Leu 290	ggt Gly	cag Gln	gtt Val	gac Asp	999 Gly 295	ggc Gly	atg Met	tcc Ser	ggt Gly	ccg Pro 300	Ala	tct Ser	tac Tyr	gcc Ala	912
aat Asn	tac Tyr	gtt Val	gtc Val	agg Arg	ggc Gly	gaa Glu	aat Asn	ctc Leu	gta Val	acc Thr	gcc	gtg Val	acg Thr	tac Tyr	ggt	960

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Pro Asn Ala	-	• • •	gat cgg gac go Asp Arg Asp Al		-
222 2 2	1.5	_	atc gcg gcg gc Ile Ala Ala Al 36	a Val Ile	_ =
•			agt tta cag co Ser Leu Gln An 380		
			agg cgt atg ca Arg Arg Met Hi 395		
		_	cgt cct cag ta Arg Pro Gln Ty 410		
Ala Thr Ile I			gcg gaa caa to Ala Glu Gln Se		
			ctg agt ttc as Leu Ser Phe As	n Tyr Gln	
• -			atg cac aac co Met His Asn Po 460		
			gat cec gga ca Asp Pro Gly H: 475		
70		_	aac atg aac c Asn Met Asn Le 490		_
Val Tyr Asn T			gtg gca cac g Val Ala His Va		
-		_	gag ttg ctg ca Glu Leu Leu H 52		
			ccg atg ttt ga Pro Met Phe As 540		
			tat agg gcg ad Tyr Arg Ala T 555		

atg Met	gtg Val	ggt Gly	aac Asn.	att Ile 565	cca Pro	caa Gln	ccc Pro	ctg Leu	gcg Ala 570	ccc Pro	aac Asn	gag Glu	ttt Phe	caa Gln 575	aac Asn	1728
agc Ser	aga Arg	ggc Gly	ctg Leu 580	cag Gln	ttt Phe	gac Asp	aga Arg	gcg Ala 585	gcg Ala	gcc Ala	gtg Val	gct Ala	cac His 590	gtg Val	ctg Leu	1776
gac Asp	cag Gln	tca Ser 595	acg Thr	atg Met	gaa Glu	att Ile	atc Ile 600	caa Gln	gat Asp	acg Thr	gcg Ala	ttt Phe 605	gac Asp	acg Thr	tcg Ser	1824
tac Tyr	cca Pro 610	cta Leu	ctc Leu	tgt Cys	tat Tyr	gtc Val 615	atc Ile	gaa Glu	tgc Cys	ctc Leu	att Ile 620	cac His	gga Gly	cag Gln	gaa Glu	1872
gac Asp 625	aaa Lys	ttt Phe	ttg Leu	att Ile	aat Asn 630	tct Ser	cct Pro	tta Leu	att Ile	gca Ala 635	tta Leu	acc Thr	att Ile	gaa Glu	act Thr 640	1920
tac Tyr	tgg Trp	aac Asn	aat Asn	gcc Ala 645	gga Gly	aaa Lys	ctg Leu	gcg Ala	ttt Phe 650	att Ile	aac Asn	agc Ser	ttc Phe	cct Pro 655	atg Met	1968
ctg Leu	cga Arg	ttt Phe	atc Ile 660	tgc Cys	gtt Val	cac His	ctg Leu	ggc Gly 665	aac Asn	ggt Gly	agt Ser	att Ile	tct Ser 670	aag Lys	gac Asp	2016
gtg Val	tac Tyr	gcc Ala 675	cat His	tac Tyr	cga Arg	aaa Lys	gtt Val 680	ttt Phe	ggc Gly	gaa Glu	ctc Leu	gtg Val 685	Val	ttg Leu	cag Gln	2064
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gcg Ala 705	Ser	gag Glu	ctg Leu	att Ile	aac Asn 710	tgt Cys	ctt Leu	cag Gln	gac Asp	Pro 715	Asr	ctt Leu	ttg Leu	ccg Pro	Pro 720	2160
ttt Phe	gct Ala	tac Tyr	aat Asn	gac Asp 725	Val	ttt Phe	acc Thr	aac Asn	ctg Leu 730	Leu	agg Arg	g cag g Glr	tcc Ser	Ser 735	cgg Arg	2208
cac	ccc Pro	atg Met	gta Val 740	Leu	ata Ile	ggc Gly	gac Asp	gag Glu 745	. Gly	tac Tyr	gaa Glu	a acg ı Thr	g gaa Glu 750	ı Asr	gac Asp	2256
agg Arg	gat Asp	acg Thr	Tyr	ato	aac Asn	gtc Val	aga Arg 760	Gly	aaa Lys	atg Met	g gag Glu	g gad a Asp 765) Let	a gto ı Val	ggt Gly	2304
gac Asp	ato Met	. Val	aac Asn	att Ile	tac Tyr	gag Glu 775	Thr	aga Arg	aac Asr	aac Asi	gcg n Ala 789	a Asp	cat His	gao S Ası	ggc Gly	2352
cgc Arg 785	g His	gtc Val	ctt Leu	gac Asp	gto Val	Gly	ccc Pro	ttt Phe	: aat : Asr	gaa Gli 799	ı Ası	c gaa n Gli	a caq u Gli	g cad	atg s Met 800	2400

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						atg Met										2496
						cca Pro										2544
						ctg Leu 855										2592
gag Glu 865	gct Ala	tcg Ser	gat Asp	ata Ile	cac His 870	ccc Pro	acc Thr	gtt Val	gac Asp	atg Met 875	att Ile	cga Arg	act Thr	ctt Leu	tgc Cys 880	2640
acg Thr	tcg Ser	ttt Phe	ctc Leu	acc Thr 885	tgc Cys	ccg Pro	ttt Phe	gtt Val	acc Thr 890	cag Gln	gcc Ala	tcc Ser	cgt Arg	gtt Val 895	gtg Val	2688
						caa Gln										2736
tac Tyr	gtg Val	agc Ser 915	cag Gln	act Thr	gtc Val	ctc Leu	gtt Val 920	aac Asn	gly ggg	ttc Phe	gcg Ala	gcg Ala 925	ttt Phe	gct Ala	atc Ile	2784
gca Ala	gat Asp 930	agg Arg	tct Ser	cgt Arg	gac Asp	gtt Val 935	gcc Ala	gag Glu	acc Thr	atg Met	ttt Phe 940	tac Tyr	ccg Pro	gtg Val	ccg Pro	2832
						gat Asp										2880
						acg Thr										2928
						gcc Ala										2976
aag Lys	tct Ser	cca Pro 995	atg Met	ctg Leu	gcc Ala	tac Tyr	gct Ala 1000	aac Asn	acc Thr	tgc Cys	Pro	atg Met 1005	acg Thr	ccc Pro	acg Thr	3024
ser					Ala	agc Ser 1015				Lys						3072
	Ile			Ala		cac His			His					Met		3120
gcc	gtc	cga	acc	gat	gag	gtg	ttg	gcg	gag	aac	ttg	cta	ttt	agt	gcc	3168

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agg gcc tcg acg tcc a	tg ttt tta ggg cag	cca tcg gtt atg cgt cgg 3216
Arg Ala Ser Thr Ser M	et Phe Leu Gly Gln	Pro Ser Val Met Arg Arg
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gaa gtc agg gcg gac g	ca gtc acg ttt gag	gtg aat cat gag ttg gca 3264
Glu Val Arg Ala Asp A	la Val Thr Phe Glu	Val Asn His Glu Leu Ala
1075	1080	1085
tcg ctg gac atg gcg c	tc ggt tat tct tcc	acc atc acg ccc gcc cac 3312
Ser Leu Asp Met Ala L	eu Gly Tyr Ser Ser	Thr Ile Thr Pro Ala His
1090	1095	1100
gtt gcg gcg att acc t Val Ala Ala Ile Thr S 1105 11	er Asp Met Gly Val	cac tgt cag gac atg ttt 3360 His Cys Gln Asp Met Phe 1115 1120
		agg acc ctc aac gac tac 3408 Arg Thr Leu Asn Asp Tyr 1135
gtt aaa caa aaa gcc g	ga tgc caa cga ttc	ggt ggt cct ggc cag att 3456
Val Lys Gln Lys Ala G	ly Cys Gln Arg Phe	Gly Gly Pro Gly Gln Ile
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cgt gag ccc gtc gct t	ac gtt gcg ggg gtg	ccg cac tcg gac aac ata 3504
Arg Glu Pro Val Ala T	yr Val Ala Gly Val	Pro His Ser Asp Asn Ile
1155	1160	1165
ccg ggt ctc agc cac g	ga cag ctg gcc acg	tgt gag att gtt ttg acg 3552
Pro Gly Leu Ser His G	ly Gln Leu Ala Thr	Cys Glu Ile Val Leu Thr
1170	1175	1180
	al Thr Tyr Phe Gln	acc ccc aac agt ccc cgg 3600 Thr Pro Asn Ser Pro Arg 1195 1200
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Gly Arg Ala Ser Cys V	al Ile Ser Cys Asp	Ala Tyr Asn Asn Glu Ser
1205	1210	1215
gcg gaa cgt ttg ctc t	tt gac cac tcc atc	ccg gat tct gcc tac gaa 3696
Ala Glu Arg Leu Leu P	he Asp His Ser Ile	Pro Asp Ser Ala Tyr Glu
1220	1225	1230
tac cgc act acg gtt a Tyr Arg Thr Thr Val A 1235	ac cca tgg gcg tcg sn Pro Trp Ala Ser	cag cag ggc tcc ctc gga 3744 Gln Gln Gly Ser Leu Gly
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gac gtg ctg tac aac t Asp Val Leu Tyr Asn S 1250	ca acc tcg cgc cag	gtc gca gtg cca ggg atg 3792 Val Ala Val Pro Gly Met 1260
Asp Val Leu Tyr Asn S 1250 tac agt ccg tgt cgc c	ca acc tcg cgc cag er Thr Ser Arg Gln 1255 ag ttt ttc cac aag ln Phe Phe His Lys	gtc gca gtg cca ggg atg 3792 Val Ala Val Pro Gly Met

. 1285 gga acg ccg gcg acc agc gcg acg gac ctg cag tac gtg gtg gtc aac Gly Thr Pro Ala Thr Ser Ala Thr Asp Leu Gln Tyr Val Val Val Asn gga acg gat gtg ttt cta gaa caa ccg tgc cag ttt cta caa gaa gcg Gly Thr Asp Val Phe Leu Glu Gln Pro Cys Gln Phe Leu Gln Glu Ala ttt ccc acg ctc gcc gcc agt cac agg tct ctg ctg gac gaa tat atg Phe Pro Thr Leu Ala Ala Ser His Arg Ser Leu Leu Asp Glu Tyr Met tog aat aag oto aog oad god oot gtg dad atg gga dat tat atg att Ser Asn Lys Leu Thr His Ala Pro Val His Met Gly His Tyr Met Ile gag gaa gtg gcc cct atg aaa aga cta tta aag atc gga aac aag gtc Glu Glu Val Ala Pro Met Lys Arg Leu Leu Lys Ile Gly Asn Lys Val qcc tat tag Ala Tyr <210> 45 <211> 1378 <212> PRT <213> Macaca mulatta rhadinovirus 17577 Met Glu Ala Ala Leu Glu Val Arg Pro Phe Pro Tyr Met Ala Thr Glu Ala Asn Leu Leu Arg Gln Met Lys Glu Ser Ala Ala Ser Gly Leu Phe Lys Ser Phe Gln Leu Leu Cly Lys Asp Ala Arg Glu Gly Gly Val Gln Phe Glu Gly Leu Leu Gly Val Tyr Thr Asn Val Ile Gln Phe Val Lys Phe Leu Glu Thr Ser Leu Ala Val Ala Cys Val Asn Thr Glu Phe Lys Asp Leu Lys Arg Met Thr Asp Gly Lys Ile Gln Phe Lys Val Ser Val Pro Thr Ile Ala Tyr Gly Asp Gly Arg Arg Pro Thr Lys Gln Lys Gln Tyr Ile Ile Met Lys Ala Cys Asn Lys His His Ile Gly Ala Glu Ile Glu Leu Ser Thr Asp Asp Ile Glu Leu Leu Phe Ile Asp Arg Glu Thr Pro Leu Asp Tyr Thr Glu Tyr Ala Gly Ala Val Lys Thr Ile Thr Ala Ser Leu Gln Phe Gly Val Asp Ala Leu Glu Arg Gly Leu Val Asp Thr Val Leu Asn Val Lys Leu Arg Ser Ala Pro Pro Met Phe Ile Leu

Lys Thr Leu Ser Asp Pro Val Tyr Thr Glu Arg Gly Leu Lys Lys Ala

Ser Phe Phe Leu Asp Lys Ser Asp Ile Ala Val Lys Gly Lys Gln Tyr

Val Lys Ser Asp Met Val Ser Met Phe Lys Ser Tyr Leu Met Asp Asn

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			Gly 260					265					270		
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	290		Gln			295					300				
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			Arg	325					330					335	
			Gln 340					345					350		
-		355	Ser				360					365			
	370		Lys			375					380				
385			Phe		390					395					400
			Ile	405					410					415	
			Lys 420					425					430		
_		435	Asn				440					445			
	450		Ser			455					460				
465			Leu		470					475					480
-	_		Arg Tyr	485					490					495	
			500 Ala					505					510		
		515	Leu				520					525			
	530		Pro			535					540				
545			Asn		550					555					560
			Leu	565					570					575	
			580 Thr					585					590		
		595					600					605			Glu
	610		Leu			615					620				
625			Asn		630					635					640
_				645					650					655	
			660					665					670		Gln
	_	675					680					685			Pro
	690					695					700				Pro
705					710	•			-	715					720

Phe Ala Tyr Asn Asp Val Phe Thr Asn Leu Leu Arg Gln Ser Ser Arg 730 725 His Pro Met Val Leu Ile Gly Asp Glu Gly Tyr Glu Thr Glu Asn Asp 740 745 750 Arq Asp Thr Tyr Ile Asn Val Arg Gly Lys Met Glu Asp Leu Val Gly 760 755 Asp Met Val Asn Ile Tyr Glu Thr Arg Asn Asn Ala Asp His Asp Gly 775 780 Arg His Val Leu Asp Val Gly Pro Phe Asn Glu Asn Glu Gln His Met 790 795 Ala Val Leu Glu Lys Leu Phe Tyr Tyr Val Val Leu Pro Ala Cys Thr 805 810 Asn Gly His Val Cys Gly Met Gly Val Asp Phe Asp Asn Val Ala Leu 820 825 Ala Leu Thr Tyr Asn Gly Pro Val Phe Ala Asp Val Val Asn Pro Asp 840 845 Asp Glu Ile Leu Asp His Leu Glu Asn Gly Thr Leu Arg Glu Met Leu 850 855 Glu Ala Ser Asp Ile His Pro Thr Val Asp Met Ile Arg Thr Leu Cys 870 875 Thr Ser Phe Leu Thr Cys Pro Phe Val Thr Gln Ala Ser Arg Val Val 890 885 Thr Gln Arg Asp Pro Ala Gln Leu Leu Thr Thr His Asp Asp Gly Arg 905 910 Tyr Val Ser Gln Thr Val Leu Val Asn Gly Phe Ala Ala Phe Ala Ile 920 Ala Asp Arg Ser Arg Asp Val Ala Glu Thr Met Phe Tyr Pro Val Pro 935 940 Phe Thr Lys Leu Tyr Ser Asp Pro Leu Val Ala Ala Thr Leu His Pro 950 955 Leu Val Ala Asn Tyr Val Thr Arg Leu Pro Ala Gln Arg Val Pro Val 970 975 965 Ala Phe Asn Val Pro Pro Ala Leu Met Ala Glu Tyr Glu Glu Trp His 980 985 990 Lys Ser Pro Met Leu Ala Tyr Ala Asn Thr Cys Pro Met Thr Pro Thr 995 1000 1005 Ser Leu Ser Thr Leu Ala Ser Met His Met Lys Leu Ser Ala Pro Gly 1015 1020 Phe Ile Cys His Ala Lys His Lys Ile His Pro Gly Phe Ala Met Thr 1030 1035 Ala Val Arg Thr Asp Glu Val Leu Ala Glu Asn Leu Leu Phe Ser Ala 1045 1050 1055 Arg Ala Ser Thr Ser Met Phe Leu Gly Gln Pro Ser Val Met Arg Arg 1060 1065 1070 Glu Val Arg Ala Asp Ala Val Thr Phe Glu Val Asn His Glu Leu Ala 1075 1080 1085 Ser Leu Asp Met Ala Leu Gly Tyr Ser Ser Thr Ile Thr Pro Ala His 1095 1100 Val Ala Ala Ile Thr Ser Asp Met Gly Val His Cys Gln Asp Met Phe 1105 1110 1115 Leu Met Phe Pro Gly Asp Ser Tyr Gln Asp Arg Thr Leu Asn Asp Tyr 1125 1130 1135 Val Lys Gln Lys Ala Gly Cys Gln Arg Phe Gly Gly Pro Gly Gln Ile 1140 1145 Arg Glu Pro Val Ala Tyr Val Ala Gly Val Pro His Ser Asp Asn Ile 1155 1160 1165 Pro Gly Leu Ser His Gly Gln Leu Ala Thr Cys Glu Ile Val Leu Thr 1175 1180 Pro Val Thr Ala Asp Val Thr Tyr Phe Gln Thr Pro Asn Ser Pro Arg 1195 1190 Gly Arg Ala Ser Cys Val Ile Ser Cys Asp Ala Tyr Asn Asn Glu Ser

				1205					210					215		
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Tyr	-	Thr 235	Thr	Val	Asn		Trp 1240	Ala	Ser	Gln		Gly 245	Ser	Leu	Gly	
Asp 1	Val 250	Leu	Туг	Asn		Thr 1255	Ser	Arg	Gln		Ala 1260	Val	Pro	Gly	Met	
Tyr 1265	Ser	Pro	Cys		Gln 1270	Phe	Phe	His		Asp 275	Ala	Ile	Leu		Asn .280	
Asn	Arg	Gly				Leu	Val		Glu 1290	Tyr	Ala	Ala		Leu 295	Thr	
Gly	Thr				Ser	Ala		Asp 1305	Leu	Gln	Tyr		Val 310	Val	Asn	
Gly				Phe	Leu		Gln 1320	Pro	Cys	Gln		Leu 1325	Gln	Glu	Ala	
			Leu	Ala				Arg	Ser		Leu 1340	Asp	Glu	Tyr	Met	
Ser 1345	Asn	Lys	Leu	Thr			Pro	Val		Met L355	Gly	His	Tyr		Ile .360	
		Val		Pro 1365		Lys	Arg				Ile	Gly		Lys 1375	Val	
Ala	Tyr		-	1305												
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	.> 91 !> DN															
			a mul	latta	a rha	adino	oviru	ıs 1	7577							
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1	7114			5					10					15		
gcc	gac	gag	ata	gca Ala	aat	ctt	cag	tca Ser	aag	ata Ile	gga Glv	tgc Cvs	att Tle	ttg Leu	cct Pro	96
Ala	Asp.	Giu	20	AIG	71511	Dea	0111	25	2,2	110	<i></i>	-1-	3 0			
ctc	aga	gac	gcc	cac His	cgt	ctg	cag	aat	ata	cag	gcg	ctg	ggt Gly	ctg	ggg Glv	144
цеп	Arg	35	ATG	ure	Arg	БСИ	40	ASII	110	0111	7114	45	OI,	200	<i>1</i>	
aac	ctg	tgc	tct	agg	gat	tcc	gcg	gtg	gat	ttt	att	cag	gca	tat	cac	192
Asn	Leu 50	Cys	ser	Arg	Asp	55 55	Ата	Val	АБР	PHE	60	GIII	АТА	lyi	1113	
tat	ttg	gac	aaa	tgc	act	ctc	gcc	gtg	ttg	gaa	gag	gtc	ggt	ccc	aac	240
Tyr 65	Leu	Asp	Lys	Cys	Thr 70	Leu	Ala	Val	Leu	Glu 75	Glu	Val	Gly	Pro	Asn 80	
aqt	tta	cgg	cta	acg	cgc	att	gat	ccc	atg	gac	aat	tat	caa	ata	aaa	288
Ser	Leu	Arg	Leu	Thr 85	Arg	Ile	Asp	Pro	Met 90	Asp	Asn	Tyr	Gln	Ile 95	Lys	
aac	qca	tac	caa	ccg Pro	gcc	ttc	cat	tgg	gat	aac	tac	tca	gaa	ttg	gta	336

110 1.05 100 gtt ata cca ccg gtc ttt ggg cgc aaa gat gcg acc gtc tca ctg gag Val Ile Pro Pro Val Phe Gly Arg Lys Asp Ala Thr Val Ser Leu Glu 120 tot aac ggg ttt gat gtg gtt ttc cet gcc gtg gtg cca gaa cca ctg Ser Asn Gly Phe Asp Val Val Phe Pro Ala Val Val Pro Glu Pro Leu 135 gct caa aca gtg ctt cag aag ctg ctg ctg tat aac ata tac tac aga Ala Gln Thr Val Leu Gln Lys Leu Leu Leu Tyr Asn Ile Tyr Tyr Arg 150 gtg geg gag acg acg eec ace gae gte aac eta gee gag gtg acg etg 528 Val Ala Glu Thr Thr Pro Thr Asp Val Asn Leu Ala Glu Val Thr Leu 170 tac acg acc aat atc act tac atg ggt cgc aac tac gcc ctg gac gtg 576 Tyr Thr Thr Asn Ile Thr Tyr Met Gly Arg Asn Tyr Ala Leu Asp Val 185 180 gac ecc gtt ggg teg age tea get atg egg atg etg gae gae etg tee Asp Pro Val Gly Ser Ser Ser Ala Met Arg Met Leu Asp Asp Leu Ser 200 att tac ctg tgc gtt ttg tcc gcg tta att ccg cgc ggg tgc gta agg 672 Ile Tyr Leu Cys Val Leu Ser Ala Leu Ile Pro Arg Gly Cys Val Arg 220 215 cta ctg acc tca ttg gtg cgc cac aac aaa cac gaa tta gtc gag att 720 Leu Leu Thr Ser Leu Val Arg His Asn Lys His Glu Leu Val Glu Ile 235 ttc gag ggg gtg gtg cca cct gag gta cag gcc ctg gat ctc aac aac 768 Phe Glu Gly Val Val Pro Pro Glu Val Gln Ala Leu Asp Leu Asn Asn 245 gta ago gtg goo gao gao ata aog ogo atg ggt goo oto ata aco tat 816 Val Ser Val Ala Asp Asp Ile Thr Arg Met Gly Ala Leu Ile Thr Tyr 265 260 cta cga agt ctc agt tct ata ttt aat ctg ggc cgc aga ttt cac gtt 864 Leu Arg Ser Leu Ser Ser Ile Phe Asn Leu Gly Arg Arg Phe His Val 280 tac gcg ttc tca tcg gac acg aat acc gct tcc tgt tgg tgt gca tat 912 Tyr Ala Phe Ser Ser Asp Thr Asn Thr Ala Ser Cys Trp Cys Ala Tyr 300 295 290 918 aac tag 305 <210> 47 <211> 305 <212> PRT <213> Macaca mulatta rhadinovirus 17577

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Leu Arg Asp Ala His Arg Leu Gln Asn Ile Gln Ala Leu Gly Leu Gly
                                        45
                       40
      35
Asn Leu Cys Ser Arg Asp Ser Ala Val Asp Phe Ile Gln Ala Tyr His
                    55
                                     60
   50
Tyr Leu Asp Lys Cys Thr Leu Ala Val Leu Glu Glu Val Gly Pro Asn
                               75
                 70
Ser Leu Arg Leu Thr Arg Ile Asp Pro Met Asp Asn Tyr Gln Ile Lys
                       90
              85
Asn Ala Tyr Gln Pro Ala Phe His Trp Asp Asn Tyr Ser Glu Leu Val
                                           110
                 105
         100
Val Ile Pro Pro Val Phe Gly Arg Lys Asp Ala Thr Val Ser Leu Glu
      115 120
                                       125
Ser Asn Gly Phe Asp Val Val Phe Pro Ala Val Val Pro Glu Pro Leu
                         140
                   135
Ala Gln Thr Val Leu Gln Lys Leu Leu Leu Tyr Asn Ile Tyr Tyr Arg
                                 155
                150
Val Ala Glu Thr Thr Pro Thr Asp Val Asn Leu Ala Glu Val Thr Leu
                                       175
                             170
           165
Tyr Thr Thr Asn Ile Thr Tyr Met Gly Arg Asn Tyr Ala Leu Asp Val
         180 185 190
Asp Pro Val Gly Ser Ser Ser Ala Met Arg Met Leu Asp Asp Leu Ser
                               205
                       200
   195
Ile Tyr Leu Cys Val Leu Ser Ala Leu Ile Pro Arg Gly Cys Val Arg
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   210
Leu Leu Thr Ser Leu Val Arg His Asn Lys His Glu Leu Val Glu Ile
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               230
Phe Glu Gly Val Val Pro Pro Glu Val Gln Ala Leu Asp Leu Asn Asn
             245
                            250
Val Ser Val Ala Asp Asp Ile Thr Arg Met Gly Ala Leu Ile Thr Tyr
                                    270
                         265
       260
Leu Arg Ser Leu Ser Ser Ile Phe Asn Leu Gly Arg Arg Phe His Val
      275
                     280
                                        285
Tyr Ala Phe Ser Ser Asp Thr Asn Thr Ala Ser Cys Trp Cys Ala Tyr
                   295
Asn
305
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														gtt Val		144
														att Ile		192
														gcg Ala		240
														ctc Leu 95		288
														gcg Ala		336
aac Asn	ccc Pro	atg Met 115	ggt Gly	cca Pro	aaa Lys	cat His	gtc Val 120	acg Thr	aaa Lys	cta Leu	ccg Pro	cac His 125	ccg Pro	gct Ala	att Ile	384
														agc Ser		432
														tct Ser		480
gat Asp	tct Ser	ggt Gly	cag Gln	cgc Arg 165	gca Ala	cac His	tat Tyr	Gly 999	ctg Leu 170	gcc Ala	ctg Leu	tta Leu	aag Lys	gcg Ala 175	gcc Ala	528
														caa Gln		576
														ttt Phe		624
tcg Ser	gca Ala 210	atg Met	gcg Ala	gca Ala	acc Thr	acg Thr 215	ttt Phe	tgc Cys	gga Gly	tcc Ser	aga Arg 220	ggc Gly	gtt Val	ctg Leu	tgg Trp	672
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		ccc Pro														810

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Tyr Pro Lys Thr Thr Asn Leu Ala Asn Glu Arg Ala Asp Val Val Lys
       35
                            40
Glu Ala Phe Asp Thr Glu Thr Pro Val Asp Ile Val Lys Gln Ile Val
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                       55
    50
Asn Glu Gly Leu Ala Ile Ser Lys Lys Asn Cys Val Arg Leu Ala Leu
                    70
                                       75
Tyr Leu Tyr Phe Tyr Leu Gln Tyr Val Cys Phe Ala Leu Leu Leu Thr
                                    90
Trp Gln Leu Asn Pro Tyr Met Asp Pro Pro Gly Leu Val Phe Ala Val
          100
                              105
Asn Pro Met Gly Pro Lys His Val Thr Lys Leu Pro His Pro Ala Ile
                          120
                                      125
       115
Val Ala Val Gly Cys Gly Ala Asp Ala Ile Cys Lys Asn Cys Ser Val
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                                           140
Pro Asp Ile Lys Thr Glu Leu Gly Met Val Tyr His Asn Gly Ser Ser
                  150
                                     155
Asp Ser Gly Gln Arg Ala His Tyr Gly Leu Ala Leu Leu Lys Ala Ala
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               165
Trp Leu Val Met Gly Asn Val Cys Pro Glu Pro Val Val Arg Gln Gly
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                              185
           1.80
Ala Ala Leu Leu Gly Pro Trp Asn Arg Thr Glu Trp Ser Asp Phe Lys
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                           200
Ser Ala Met Ala Ala Thr Thr Phe Cys Gly Ser Arg Gly Val Leu Trp
                                           220
                       215
Ser Pro Ile His Glu Lys Asn Leu Cys Arg Pro Thr Trp Asn Asp Val
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                                      235
Ile Asn Thr Ser Val Phe Thr Asn Glu Ser Leu Cys Pro Asn Ile Pro
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              245
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agt cag ccg gaa tcg gtt caa gtt tct cca ttt tat cgc gta att aca
Ser Gln Pro Glu Ser Val Gln Val Ser Pro Phe Tyr Arg Val Ile Thr
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gcg ttg g Ala Leu V 50	gta tgg Val Trp	tac gt Tyr Va	g atg 1 Met 55	cgg Arg	agg Arg	gtg Val	tgt Cys	tgt Cys 60	aag Lys	gly aaa	cgc Arg	gtt Val	192
gtt gcc g Val Ala A 65	gat tcg Asp Ser	Cys Ai	gc gac g Asp '0	ccg Pro	cgt Arg	caa Gln	ccc Pro 75	gcg Ala	tat Tyr	gag Glu	atg Met	ttg Leu 80	240
aat gtt a Asn Val A	agg ttg Arg Leu	cgt co Arg Pi 85	c cac o His	gga Gly	acc Thr	aat Asn 90	cca Pro	tag					276
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Ser Gln F		Ser V	al Gln	Val			Phe	Tyr	Arg				
Lys Pro E	20 Pro Val	Met G	ly Leu		25 Phe	Cys	Va1	Ala		30 Cys	Val	Ile	
Ala Leu V	35 Val Trp	Tyr Va	al Met	40 Arg	Arg	Val	Cys	Cys	45 Lys	Gly	Arg	Val	
50			55					60					
Val Ala <i>I</i> 65		•	70				75	Ата	тут	Giu	мес	80	
Asn Val A	Arg Leu	Arg P	ro His	Gly	Thr	Asn 90	Pro						
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	cacaa	,											
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1		5				10					15	•	
tct gac	aaa tca	act a	gt ttt	tta	ctt	aat	ttg	aag	gad	gcc	cac	gaa	96
Ser Asp	Lys Ser 20		er bue	: Leu	ьеи 25		nen	гъл	, wat	30		, Giu	
aag atg	ctt aac	gtg g	tg aad	tac	gta	tgt	ccg	gat	cat	. aaa	ı gat	gat	144
Lys Met	Leu Asn	Val V	al Asr	Tyr 40	Val	Сув	Pro	Asp	His 45	r PA	as Asp	Asp	
	35												100
ttt aac	ttg caa	gac a	ct gto	gtg	gcg	tgc	ccg	tgo	: tac	c cgc	ctg	y cat	192

Phe	Asn 50	Leu	Gln	Asp	Thr	Val 55		Ala	Cys	Pro	Cys 60	Tyr	Arg	Leu	His	
		gcc Ala			_			-	_	-		_				240
		ttg Leu	_						_				_		_	288
_		gct Ala		_	-					_	_		_	-		336
	_	gat Asp 115	-		_	_	_	_	_			_		_		384
	-	aag Lys	-	_			-		_		-					432
		gac Asp													aat Asn 160	. 480
	_	gtg Val								_	-	,				528
_	-	cta Leu				_	-		_		-		_	_	_	576
		att Ile 195									_		_		-	624
	_	gcç Ala					_	_		_				_		672
_		gtg Val					_	_	_			_				720
		gac Asp	_						-				_	_		768
	~ _	aaa Lys		_			-					~	_		_	816
	_	ttt Phe 275			_									-		864
_		gat Asp		_								_			_	912

295 300 290

tgc att cca cta aaa gac gga ggt cac acg tac tgc gcg aaa caa aaa Cys Ile Pro Leu Lys Asp Gly Gly His Thr Tyr Cys Ala Lys Gln Lys 315 310

acc atg teg gac gac gtg ett gte gee gee gte atg gee cae tac atg 1008 Thr Met Ser Asp Asp Val Leu Val Ala Ala Val Met Ala His Tyr Met 330 325

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<210> 53

<211> 348

<212> PRT

<213> Macaca mulatta rhadinovirus 17577

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Ser Asp Lys Ser Thr Ser Phe Leu Leu Asn Leu Lys Asp Ala His Glu 30 20 25

Lys Met Leu Asn Val Val Asn Tyr Val Cys Pro Asp His Lys Asp Asp 4.5 40 35

Phe Asn Leu Gln Asp Thr Val Val Ala Cys Pro Cys Tyr Arg Leu His 55

Ile Pro Ala Tyr Ile Thr Ile Asp Glu Thr Val Arg Ser Thr Thr Asn 75 70 Leu Phe Leu Glu Gly Ala Phe Ser Thr Glu Leu Met Gly Asp Ala Ala

90 Thr Ser Ala Gln Ser Met His Lys Ile Val Ser Asp Ser Ser Leu Ser

100 Gln Leu Asp Leu Cys Arg Val Lys Ser Thr Ser Gln Asp Ile Gln Gly

120 Ala Met Lys Pro Cys Leu His Val Tyr Ile Asp Pro Ala Tyr Thr Asn 140 135

Asn Thr Asp Ala Ser Gly Thr Gly Ile Gly Ala Val Ile Ala Val Asn 155 150

His Lys Val Ile Lys Cys Ile Leu Leu Gly Val Glu His Phe Phe Leu 170 175 165

Arg Asp Leu Thr Gly Thr Ala Ala Tyr Gln Ile Ala Ser Cys Ala Ala 185 180

Ala Leu Ile Arg Ala Ile Val Thr Leu His Pro Gln Ile Thr His Val 200 205

Asn Val Ala Val Glu Gly Asn Ser Ser Gln Asp Ala Gly Val Ala Ile 220 215 210

Ala Thr Val Leu Asn Glu Ile Cys Ser Val Pro Leu Ser Phe Leu His 235 230

His Val Asp Lys Asn Thr Leu Ile Arg Ser Pro Ile Tyr Met Leu Gly 250 245

Pro Glu Lys Ala Lys Ala Phe Glu Ser Phe Ile Tyr Ala Leu Asn Ser 265 260

Gly Thr Phe Ser Ala Ser Gln Thr Val Val Ser His Thr Ile Lys Leu 280

Ser Phe Asp Pro Val Ala Tyr Leu Ile Asp Gln Ile Lys Ala Ile Arg 300 295 Cys Ile Pro Leu Lys Asp Gly Gly His Thr Tyr Cys Ala Lys Gln Lys 315

310

Thr Met Ser Asp Asp Val Leu Val Ala Val Wet Ala His Tyr Met
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340 - 340 - 345

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cag tgt cag gcg ttc ttt cac cgt ccc att aga gat cta att tca tct 96
Gln Cys Gln Ala Phe Phe His Arg Pro Ile Arg Asp Leu Ile Ser Ser
20 25 30

gga gct gac gct tta aac cac ttt agc cta tct gaa tca gac gga cat 144 Gly Ala Asp Ala Leu Asn His Phe Ser Leu Ser Glu Ser Asp Gly His 35 40 45

aaa ttg gaa cgg att gtt ctt ctg ctt gac ctg gtg ggg aca gaa tgt 192 Lys Leu Glu Arg Ile Val Leu Leu Leu Asp Leu Val Gly Thr Glu Cys 50 55 60

ctc tct tat acc acg atc gct gca aag aat gtc aaa tga 231 Leu Ser Tyr Thr Thr Ile Ala Ala Lys Asn Val Lys 75

<210> 55 <211> 76 <212> PRT

<213> Macaca mulatta rhadinovirus 17577

<400> 55

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35 40 45 Lys Leu Glu Arg Ile Val Leu Leu Leu Asp Leu Val Gly Thr Glu Cys 50 55 60

Leu Ser Tyr Thr Thr Ile Ala Ala Lys Asn Val Lys 65 70 75

<210> 56

<211> 654

<212> DNA

<213> Macaca mulatta rhadinovirus 17577

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gtc aac agc co Val Asn Ser Pr	ca ata tgt cga co Ile Cys Arg	ttt cat aac Phe His Asn 25	gtc tct aac (Val Ser Asn)	tta tac cag 96 Leu Tyr Gln 30	
tgt ttg gat to Cys Leu Asp Cy 35	gt aag cgc tat vs Lys Arg Ty:	cac gta tgc His Val Cys 40	gac ggg gga Asp Gly Gly 45	cgc aac tgc. 144 Arg Asn Cys	
gtg atc gtg ta Val Ile Val Ty 50	ac act cgc gaa /r Thr Arg Gli 55	ı Asn Leu Val	tgt gat tta Cys Asp Leu 60	acg gga aac 192 Thr Gly Asn	
tgc gtt ttg ga Cys Val Leu As 65	at aat gtg cag sp Asn Val Gli 70	g gac gta tgt n Asp Val Cys	tcg tac ggt Ser Tyr Gly 75	cct cca gaa 240 Pro Pro Glu 80	J
cgc cgc gta co Arg Arg Val Pi	cc gac gcc tto co Asp Ala Pho 85	atc gat ccg E Ile Asp Pro 90	Leu Val Ser	cac ggc acg 288 His Gly Thr 95	į
agg gaa tgt ct Arg Glu Cys Le	tt aaa agc ga eu Lys Ser As 00	ata ctg agg o Ile Leu Arg 105	g tac ttt gag g Tyr Phe Glu	acg gtc ggt 336 Thr Val Gly 110	;
gtg aaa tot ga Val Lys Ser G 115	ag gca tat to lu Ala Tyr Se	acc gtt gtc Thr Val Val	aag aat gga Lys Asn Gly 125	caa ttg aat 384 Gln Leu Asn	ł
ggc atc ata gg Gly Ile Ile G 130	gt aga tta at ly Arg Leu Il 13	e Asp Ala Thr	g ttt aac gag c Phe Asn Glu 140	tgc ctt ccg 432 Cys Leu Pro	2
gta atg agc g Val Met Ser A 145	ac ggc gaa gg sp Gly Glu Gl 150	t ggc aga gac y Gly Arg Asp	c ctc gcg gcg o Leu Ala Ala 155	agc att tac 480 Ser Ile Tyr 160)
atc cac ata a Ile His Ile I	tt atc tcc at le Ile Ser Il 165	a tac tcc act e Tyr Ser Thi 170	r Lys Thr Val	tat gat aat 528 Tyr Asp Asn 175	В
Leu Leu Phe L	aa tgt acg ag ys Cys Thr Ar 80	a aat aaa aaa g Asn Lys Lys 185	a tac gac cac s Tyr Asp His	att gta aaa 570 Ile Val Lys 190	6
act atc aga g Thr Ile Arg A 195	cg caa tgg at la Gln Trp Me	g cgc atg gto t Arg Met Val 200	c tca acc ggc l Ser Thr Gly 205	gat ccg tcg 62 Asp Pro Ser	4
cgg gtc agt g Arg Val Ser A 210	cg acg ggt tg la Thr Gly Cy 21	s Phe Thr	a	65	4

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            2.0
Cys Leu Asp Cys Lys Arg Tyr His Val Cys Asp Gly Gly Arg Asn Cys
                           40
       35
Val Ile Val Tyr Thr Arg Glu Asn Leu Val Cys Asp Leu Thr Gly Asn
                                           60
                        55
    50
Cys Val Leu Asp Asn Val Gln Asp Val Cys Ser Tyr Gly Pro Pro Glu
                                       75
Arg Arg Val Pro Asp Ala Phe Ile Asp Pro Leu Val Ser His Gly Thr
                                   90
                85
Arg Glu Cys Leu Lys Ser Asp Ile Leu Arg Tyr Phe Glu Thr Val Gly
                           105
Val Lys Ser Glu Ala Tyr Ser Thr Val Val Lys Asn Gly Gln Leu Asn
                           120
                                        125
      115
Gly Ile Ile Gly Arg Leu Ile Asp Ala Thr Phe Asn Glu Cys Leu Pro
                      135
                                          140
    130
Val Met Ser Asp Gly Glu Gly Gly Arg Asp Leu Ala Ala Ser Ile Tyr
                                       155
                   150
Ile His Ile Ile Ile Ser Ile Tyr Ser Thr Lys Thr Val Tyr Asp Asn
                                170
               165
Leu Leu Phe Lys Cys Thr Arg Asn Lys Lys Tyr Asp His Ile Val Lys
                                                190
                       185
         180
Thr Ile Arg Ala Gln Trp Met Arg Met Val Ser Thr Gly Asp Pro Ser
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                                              205
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Arg Val Ser Ala Thr Gly Cys Phe Thr
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ggg ttg ttt cac gtg ata ctt ccg cga ggg ttt atc ctc gcg aac aat
                                                                 96
Gly Leu Phe His Val Ile Leu Pro Arg Gly Phe Ile Leu Ala Asn Asn
             20
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Ile Thr Cys Gly Glu Arg Gln Arg Phe Phe Ala His Thr Trp Phe Ala
         35
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<210> 57

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caa Gln 65	aac Asn	acc Thr	gac Asp	ccg Pro	ggc Gly 70	cgc Arg	ggg Gly	gac Asp	ggt Gly	ccg Pro 75	tcc Ser	gly ggg	ccg Pro	tgg Trp	tcc Ser 80	240
gga Gly	ctg Leu	gcg Ala	att Ile	agt Ser 85	ctg Leu	cct Pro	ctg Leu	ttt Phe	acc Thr 90	aca Thr	aat Asn	gga Gly	aaa Lys	ttt Phe 95	cat His	288
ccg Pro	ttt Phe	gat Asp	gta Val 100	gtt Val	ata Ile	ctc Leu	aag Lys	gcc Ala 105	gat Asp	acg Thr	cct Pro	gac Asp	tct Ser 110	gga Gly	agc Ser	336
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aac Asn	gca Ala 130	atg Met	cga Arg	ggt Gly	tta Leu	aaa Lys 135	gat Asp	aaa Lys	gtt Val	tcg Ser	caa Gln 140	tgt Cys	acc Thr	gat Asp	gcc Ala	432
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tgg gta aag gcg gcg o Trp Val Lys Ala Ala 1 355			
gat gtt ttg gga tta t Asp Val Leu Gly Leu 3 370	0.00		
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ccg ggg ggc ttt gtt a Pro Gly Gly Phe Val I 435		=	= -
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<213> Macaca mulatta rhadinovirus 17577

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att Ile	cta Leu	gat. Asp	ttt Phe	tgt Cys 85	gcc Ala	acg Thr	gaa Glu	tct Ser	gtc Val 90	gcc Ala	ata Ile	agg Arg	gac Asp	gtg Val 95	cct Pro	288
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Thr Cys Lys Ile Cys Thr Gly Ala His Val His Val Asn Pro Tyr Arg 215 220 Gly Tyr Thr Pro Pro Asp Ser Gln Gly Thr Ser Pro Ser Cys Pro Cys 230 235 Leu Ile Ser Cys Gly Ala Arg Arg Ala Ala Asp Val Leu Val Thr Gly 250 245 His Val Asn Leu Leu Gly Leu Leu Phe Asp Pro Lys Ala Ser Pro Lys 270 260 265 Val Thr Lys Leu Arg Leu Lys Arg Asn Pro Arg Pro Val Pro Ile Glu 285 275 Asp Ala Met Ser Gly Val Thr Ala Glu Gly Thr Glu Val Gln Pro Thr 300 2.95 Ser Leu Pro Trp Ala Leu Ile Arg Leu Pro Asp Leu Ala Ser Arg Val 315 310 Met Leu Tyr Gly Cys Gln Asn Leu Lys Ser Ile Cys Leu Arg Ser Tyr 330 <210> 62 <211> 984 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(984) atq ttq tta acc agc tat cgc gaa cgt ctt caa aat aac ttg cgc gtg Met Leu Leu Thr Ser Tyr Arg Glu Arg Leu Gln Asn Asn Leu Arg Val gtc acg gac ggt ggt tgc gaa aac tgg ttt cgg caa ccg ccc gtt att Val Thr Asp Gly Gly Cys Glu Asn Trp Phe Arg Gln Pro Pro Val Ile 20 25 ata toq qqc aac gac aag acc gaa cga atg gcc cac cca tgc ttg gga 144 Ile Ser Gly Asn Asp Lys Thr Glu Arg Met Ala His Pro Cys Leu Gly 40 35 gtt att cac gcg gtt aat gca tat agt tct gtt tta gac gat tat ctt 192 Val Ile His Ala Val Asn Ala Tyr Ser Ser Val Leu Asp Asp Tyr Leu 55 caa acq tac eqc aga qtt caa qaa ecc atg eeg gee eet acg ttg gga 240 Gln Thr Tyr Arg Arg Val Gln Glu Pro Met Pro Ala Pro Thr Leu Gly 70 aag ccc cga att tet age cac get acg ttg ccc cgg tta acc gag gaa 288 Lys Pro Arg Ile Ser Ser His Ala Thr Leu Pro Arg Leu Thr Glu Glu 336 ctc aca aac tac ctt aaa caa aca tgt tgt cgg gtc caa atg gca aac Leu Thr Asn Tyr Leu Lys Gln Thr Cys Cys Arg Val Gln Met Ala Asn 105 100 gee aag gae cag tae atg gaa tae caa teg gee caa egg ace cae gaa Ala Lys Asp Gln Tyr Met Glu Tyr Gln Ser Ala Gln Arg Thr His Glu

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gta Val 305	agc Ser	aga Arg	tgc Cys	ttt Phe	tta Leu 310	agt Ser	tct Ser	ggc Gly	agc Ser	cgt Arg 315	ata Ile	gca Ala	tca Ser	cgc Arg	gac Asp 320	960
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		Gly 999											240
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_		tgg Trp										_	480
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	_	cgc Arg				_	_						576
		gcc Ala 195											624
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<213> Macaca mulatta rhadinovirus 17577

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ccc att ccc aag gtt cag act gac gtc gac aga aca gca tcg tcc cat 144
Pro Ile Pro Lys Val Gln Thr Asp Val Asp Arg Thr Ala Ser Ser His
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ata acc gtc att aaa aca cgt aag acg atc gcc caa ctg aag ata cct 192
Ile Thr Val Ile Lys Thr Arg Lys Thr Ile Ala Gln Leu Lys Ile Pro
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Asn Asn Trp Gly Gln Cys Ser His Gln Ala Thr Asp Trp Thr Ala Val

65 70 80

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tgc gtt aag cat ttt ggc agc cgg cgt gag ttt ttt tac gag tgc att 336

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_	_			-	_		aca Thr			_						480
							ctg Leu									528
_		_	_				aat Asn		_			-		-	_	576
			_			_	ata Ile 200	_		_	_					624
							cta Leu									672
						_	gtg Val									720
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		_	_	_			tac Tyr		_	_	_	_				816
					_		aat Asn 280	_	_		_	_	_			864
							tcg Ser						-			912
_						_	cca Pro	_	_				_			960
		-		-			gaa Glu		_			_			_	1008
-	_				_		gtc Val		_					_		1056

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215

Thr Pro Trp Thr Gly Val Met Val Thr Ser Lys Leu Gly Phe Val Gln 230 235 His Thr Tyr His Phe Lys Ala Pro Ala Arg Phe Ile Cys Lys His Ile 255 250 245 Tyr Arg Pro Ser Cys Leu Leu Tyr Arg Cys Leu Leu Ser Cys Ala Gly 270 260 265 Gly Pro Gln Ala His Met Leu Asn Gln Pro Phe Gln Ile Thr Pro Gln 285 280 Leu Gly Leu Thr Ile Asp Ile Ser Ser Leu Gly Tyr Ser Leu Leu Ala 300 295 290 Cys Leu Glu Lys Tyr Leu Gln Pro Ala Asp Pro Phe Pro Gln Gln Gly 310 315 Ala Leu Ala Asp Ala Ser Ser Glu Ser Ala His Pro Leu Phe Tyr Leu 330 335 325 Arg Cys Met Val Pro Arg Val Val Ile Ala Glu Ile Phe Ser Val Ala 350 345 340 Trp Asp Val Pro Leu Asp Leu Gly Ile Asp Ser Ser Gly His Ala Pro 355 365 360 Ala Ile Pro Leu Arg Glu Ala Tyr Arg Arg Phe Phe Ala Asn Gln Cys 380 375 Ser Leu Tyr Arg Ala Gln Tyr Lys Glu Asp Ala Leu Glu Asn Ala Ser 395 385 Ser Arg Leu Cys Asn Ser Lys Leu Lys Leu Val Leu Gln Lys Leu Leu 410 415 405 Val Arg Asp Tyr Phe Ser His Cys Gly Asn Cys Gly Asp His Gly Phe 425 420 Phe Leu Arg 435

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				Leu						gat Asp						336
	_		_	_	-	_	_			aac Asn						384
_	_	-		-				_		tca Ser					_	432
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_				Phe			_	-		atc Ile 235		_			_	720
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Val	Lys	Lys	Cys	Met 325	Ile	His	Asn	Ser	Thr 330	Ala	Pro	Ser	Asp	Val 335	Tyr	
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	_	agc Ser 355	_						-							1104
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_		act Thr			_			_								1200
		aga Arg			_			_	_	_	_	_	_			1248
		gcc Ala														1296
		ccg Pro 435							_				_		_	1344
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Ala Leu Ile Glu Gln Val Thr Arg Gly Gln Asn Ile Asn Pro Leu Trp
              120
                               125
      115
Asp Ala Leu Arg Asp Gly Ile Ile Ser Ser Ser Lys Phe His Trp Ala
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                            140
Ile Lys Gln Gln Asn Ser Ser Lys Lys Ile Phe Asn Pro Trp Pro Ile
                                 155
               150
Val Asn Asn His Phe Val Ala Gly Pro Leu Ala Phe Gly Leu Arg Cys
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                              170
             165
Glu Glu Val Val Lys Lys Ile Leu Ala Thr Leu Leu His Pro Gly Glu
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          180
Ala His Cys Glu Asn Tyr Gly Phe Met Gln Ser Pro Leu Asn Gly Val
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Phe Gly Val Ser Leu Asp Phe Gly Ile Asn Val Arg Ser Asp Pro Lys
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Asp Gly Leu Glu Phe His Pro Asp Cys Lys Ile Tyr Glu Ile Lys Cys
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Arg Phe Lys Tyr Thr Phe Ser Lys Met Glu Cys Asp Pro Ile Tyr Ala
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Ala Tyr Ala Lys Leu Tyr Gln Lys Pro Ser Met Gln Thr Leu Lys Gly
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                           265
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Phe Leu Tyr Ser Ile Ser Lys Pro Ala Ile Glu Phe Val Gly Glu Asp
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                       280
Arg Leu Pro Ser Glu Ser Asp Tyr Leu Val Ala Tyr Asp Lys Glu Trp
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                             300
Glu Val Cys Pro Arg Lys Lys Arg Arg Leu Thr Ala Val His His Leu
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Val Lys Lys Cys Met Ile His Asn Ser Thr Ala Pro Ser Asp Val Tyr
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His Leu Ser Ala Asn Leu Phe Ile Asn Val Arg His Pro Tyr Tyr
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                      360
      355
Gln Val Leu Leu Gln Ser Leu Val Val Gln Glu Tyr Ile Ser Leu Ser
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Lys Gly Thr Lys Asn Leu Gly Thr Gln Lys Asn Phe Ile Ala Thr Gly
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Phe Phe Arg Lys Arg Gln Phe Gln Asp Pro Ser Cys Cys Thr Ile Gly
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Glu Phe Ala Pro Leu Asp Pro His Val Glu Ile Pro Thr Leu Leu Ile
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Val Thr Pro Val Tyr Phe Pro Ser Val Ala Lys His Gln Leu Val Lys
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Gln Ala Thr Glu Phe Trp Ala Ala Ser Ala Arg Glu Ala Phe Pro Glu
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gtg aag ggg Val Lys Gly	gag ccc Glu Pro 20	ata gat Ile Asp	Val Se	cc aaa er Lys 25	gaa ttc Glu Phe	Asp P	ct att ro Ile 30	ata Ile	96
gga gaa gaa Gly Glu Glu 35	Ser Ile	gtc ttg Val Leu	tta ac Leu Th 40	cg gca hr Ala	gat ggg Asp Gly	act g Thr A 45	cc ccc la Pro	gcg Ala	144
gcg ctg tac Ala Leu Tyr 50	aaa ccc Lys Pro	aaa acc Lys Thr 55	aag co Lys Pi	ca tcc ro Ser	aaa cat Lys His 60	aaa a Lys A	ac aat sn Asn	aaa Lys	192
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cta ctg gt Leu Leu Va	t att ctt l Ile Leu 20	gga ctt i Gly Lei	atg t 1 Met E	tt ata Phe Ile 25	atg tc Met Se	a gcg r Ala	gta gt Val Va 30	g cca l Pro	96
ctg acc gc Leu Thr Al	a Thr Phe	ccg gga	a ctt g / Leu C 40	gga ttt Gly Phe	ccg tg Pro Cy	c tac s Tyr 45	ttt aa Phe As	c acg n Thr	144

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gtt Val 145	ctg Leu	tcg Ser	tac Tyr	aaa Lys	cat His 150	ata Ile	ctt Leu	tta Leu	gcc Ala	tcg Ser 155	ttt Phe	gta Val	tac Tyr	tgt Cys	ata Ile 160	480
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agc Ser	ttg Leu	cta Leu 195	gac Asp	acc Thr	ctg Leu	ttg Leu	cgt Arg 200	tac Tyr	gga Gly	aaa Lys	ccg Pro	att Ile 205	ggc	gcc Ala	aat Asn	624
ctt Leu	tat Tyr 210	ctg Leu	tcc Ser	tta Leu	ata Ile	gcc Ala 215	atg Met	gag Glu	atg Met	tta Leu	gta Val 220	ttc Phe	tcc Ser	ctc Leu	gga Gly	672
Thr 225	Met	Met	Ala	Ile	Gly 230	Asn	Ser	Phe	Tyr	Met 235		Val	Ser	Asp	11e 240	720
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atc Il∈	aac Asn	aca Thr	gaa Glu 260	cta Leu	ttt Phe	cta Leu	gta Val	aag Lys 265	Tyr	cta Leu	aag Lys	cac His	cag Glr 270	ı Ile	gga Gly	816
tto Phe	tac Tyr	gtt Val 275	Gly	gtt Val	ttt Phe	gtc Val	agt Ser 280	Tyr	ctg Leu	att Ile	ctg Lev	ctt Lev 285	ı Leı	Pro	gtc Val	864

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Leu	Val 50	Asn	Tyr	Ser	Ala	Leu 55		Leu	Thr	Val	Arg 60	Ser	Ser	Ala	Lys	
His 65	Leu	Thr	Pro	Thr	Leu 70	Phe	Leu	Glu	Ala	Pro 75	Glu	Met	Phe	Val	Tyr 80	
Ile	Ser	Trp	Ala	Phe 85	Leu	Val	Asp	Gly	Tyr 90	Leu	Leu	Cys	Tyr	Tyr 95	Ala	
Trp	Ala	Ile	Leu 100		Ile	Phe	Lys	Ala 105	Lys	Arg	Val	His	Ala 110	Thr	Thr	
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145 His	Phe	Cys	Leu		150 Phe	Thr	His	Val	Gln 170			Ile	Ser	Cys	Asn	
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Ser	Leu		180 Asp	Thr	Leu	Leu		185 Tyr	Gly	Lys	Pro	Ile 205	Gly		Asn	
Leu		195 Leu	Ser	Leu	Ile		200 Met	Glu	Met	Leu		Phe		Leu	Gly	
Thr 225	210 Met	Met	Ala	Ile	Gly 230	215 Asn	Ser	Phe	Tyr	Met 235			Ser	Asp	11e 240	

Val Phe Gly Ser Ile Asn Leu Phe Phe Val Leu Thr Ile Ala Trp Tyr 245 250 Ile Asn Thr Glu Leu Phe Leu Val Lys Tyr Leu Lys His Gln Ile Gly 265 270 260 Phe Tyr Val Gly Val Phe Val Ser Tyr Leu Ile Leu Leu Pro Val 285 275 280 Val Arg Tyr Asp Lys Val Phe Ile Ser Ala Ser Leu His Lys Val Ile 300 295 Ala Val Asn Ile Ser Met Ile Pro Ile Thr Cys Ile Leu Ala Ile Ile 310 315 Leu Arg Ile Ile Arg Asn Asp Trp Lys Trp Cys Ala Lys Ser Pro Glu 325 330 Tyr Ala Pro Leu Pro Gln Gly Pro Lys Glu Lys Thr Thr Lys Val Lys 345 350 Tyr Ser Pro Glu Leu Asn Ala Leu Tyr Glu Thr Glu Glu Asp Val Ser 360 355 Asp Tyr Glu Asp Ala Tyr Pro Lys Tyr Ile 375

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115 120 125

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gly aaa	tta Leu	tac Tyr	tat Tyr 180	agg Arg	gca Ala	tcg Ser	agt Ser	gcc Ala 185	acg Thr	caa Gln	tgc Cys	cga Arg	aaa Lys 190	agg Arg	gcg Ala	576
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gcc Ala	att Ile	gat Asp 435	aaa Lys	aca Thr	aca Thr	gta Val	ttc Phe 440	atc Ile	aag Lys	gct Ala	ccc Pro	cag Gln 445	ctc Leu	agc Ser	gca Ala	1344
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1 Leu	His	Ser	Thr	5	Cl.,	T) ====	~7							15		
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	. Val	35 Thr	20 Cys	Gln	Gln	Cys Ser	Gly 40	25 Ile	Leu Gln	Gly	Asp	Trp Ala 45 Asn	Gln 30 Ala	Val Tyr	His	
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205 200 195 Glu Ile Gln Val Ala Gly Gln Lys Tyr Thr Leu Ser Ile Ala Thr Ala 220 215 Thr Phe His Val Leu Trp Val Asp Glu Ala Cys Met Trp Asn Gly Ala 235 230 Leu Ala Glu Phe Phe Arg Ala Leu His Asn Lys Leu Phe Gly Asp Arg 250 245 Glu Gly Val Ala Pro Thr Leu Thr Tyr Val Cys Pro Gly Ala Thr Pro 265 270 260 Glu Gly Thr Pro Phe Pro Pro Tyr Phe Ser Ala Phe Pro His Leu Pro 285 280 Leu Val Phe Gly Arg Pro Arg Arg Leu Asp Val Thr Ala Val Gln Glu 300 295 Leu Pro Lys Ala Gln Ile Ala Val His Trp Pro Pro Phe Lys Asp Ser 315 310 Ile Leu Gly Asp Gln Leu Leu Ile Pro Gly Ile Ser Pro Lys Lys Pro 335 330 325 Gly Thr Val Pro Val Arg Trp Pro Leu Trp Val Glu Asp Val Asn Leu 345 340 Ser Leu Cys Glu Thr Thr Glu Ser Val Ala Arg Ile Val Asp Pro His 360 365 355 Ser Ile Val Ile Ile Lys Phe Ser Ser Leu Leu Cys Gln His Leu Lys 380 375 Cys His Arg Ala Phe Val Lys Asn Glu Leu Glu Tyr Ile Ala Thr Ile 395 390 Cys Ser Ser Asp Leu Arg Leu Phe Ile Gln Glu Glu Tyr Asn Arg Leu 410 405 Leu Ala Thr Ile Phe Thr Trp Ala Ala Ala Ser Gly Tyr Thr Trp Ala 430 425 420 Ala Ile Asp Lys Thr Thr Val Phe Ile Lys Ala Pro Gln Leu Ser Ala 440 445 435 Ala Val Ser Gly Phe Cys Pro Ser Leu Asn Ser Cys Arg Arg Lys Gln 455 Cys Tyr Glu Gly 465 <210> 78 <211> 612 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(612) <400> 78 atg tta cga agg tta aaa ata aca gtt cat ttc ctt tca cag gaa cag 48 Met Leu Arg Arg Leu Lys Ile Thr Val His Phe Leu Ser Gln Glu Gln 10 5 caa aag gtc gtg acc cgt ctt gag gcg cat ttg gga ctt ccc gta cag 96 Gln Lys Val Val Thr Arg Leu Glu Ala His Leu Gly Leu Pro Val Gln 25 gaa act too cac oog oot gao tgg oto aag tgt gag gto tgo too gog 144 Glu Thr Ser His Pro Pro Asp Trp Leu Lys Cys Glu Val Cys Ser Ala 40 tcc gtg ttt tta aaa ata cca gcc ggg gtt ttg tat gcc gga ctc gca 192

Ser	Val 50	Phe	Leu	Lys	Ile	Pro 55	Ala	Gly	Val	Leu	Tyr 60	Ala	Gly	Leu	Ala	
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gta Val	gaa Glu	ggc Gly	gcg Ala	acg Thr 85	ttg Leu	ttg Leu	ctt Leu	aac Asn	aac Asn 90	tca Ser	gtg Val	tta Leu	ccg Pro	att Ile 95	999 Gly	288
gcg Ala	ctg Leu	gcg Ala	ggt Gly 100	atc Ile	tta Leu	ccc Pro	acc Thr	ctt Leu 105	ttt Phe	gcc Ala	aac Asn	agg Arg	cgg Arg 110	tgt Cys	gtt Val	336
aat Asn	ttt Phe	tgg Trp 115	ctg Leu	ctg Leu	cca Pro	cgc Arg	gcg Ala 120	tgg Trp	gta Val	aaa Lys	tcg Ser	gcg Ala 125	ccc Pro	ata Ile	tgc Cys	384
cct Pro	ccc Pro 130	cta Leu	ccg Pro	att Ile	gac Asp	tgt Cys 135	gtt Val	acg Thr	cct Pro	cca Pro	cag Gln 140	ttt Phe	gtc Val	gtg Val	aca Thr	432
aag Lys 145	cgt Arg	gga Gly	cca Pro	atc Ile	tgc Cys 150	tgg Trp	tac Tyr	aag Lys	gaa Glu	tgg Trp 155	ccg Pro	tta Leu	ccg Pro	gtt Val	gac Asp 160	480
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65	_				70					75					80 Gly	
				85					90					95		
			100					105					110		Val	
Asn	Phe	Trp	Leu	Leu	Pro	Arg	Ala	Trp	val	гуѕ	ser	ата	Pro	тте	: Cys	

125 120 Pro Pro Leu Pro Ile Asp Cys Val Thr Pro Pro Gln Phe Val Val Thr 140 135 Lys Arg Gly Pro Ile Cys Trp Tyr Lys Glu Trp Pro Leu Pro Val Asp 155 150 Val Asp Phe Met Tyr Tyr Leu Gln Glu Ala Leu Cys Val Phe Ser Val 175 170 165 Val Ser Asn Gly Glu Gly Thr Glu Ser His Ala Asp Asn Ile Arg Gln 185 Leu Glu Lys Phe Glu Lys Val Leu Cys Leu Phe 195

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aac Asn 225	gtt Val	gga Gly	gaa Glu	gaa Glu	gag Glu 230	ggc Gly	cac His	gcg Ala	gag Glu	act Thr 235	ttt Phe	aac Asn	ata Ile	ttt Phe	tat Tyr 240	720
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Leu Lys Thr Arg Trp Arg Gly Val Glu His Leu Thr Pro Glu Phe Lys
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Arg Ser Thr Phe Glu Ser Trp Ala Arg Thr Val Arg Leu Thr Val Asp
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105
110

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agc Ser	ttt Phe 450	ttt Phe	tca Ser	caa Gln	tac Tyr	gtc Val 455	cct Pro	ccg Pro	ttt Phe	atg Met	gag Glu 460	atg Met	ctc Leu	aaa Lys	gag Glu	1392
Leu 465	Thr	Ser	ctg Leu	Trp	Glu 470	Gly	Glu	Met	Phe	Gln 475	Thr	Tyr	Asn	Leu	Thr 480	1440
Pro	Val	Val	gac Asp	Asn 485	Gln	Gly	Gln	Arg	Thr 490	Ser	Ile	Ala	Tyr	Ser 495	Gln	1488
Āsp	Thr	Val	tcc Ser 500	Ile	Leu	Leu	Gly	Pro 505	Phe	Thr	Tyr	Ile	Ile 510	Ala	Lys	1536
Leu	Thr	His 515	Met	Asp	Leu	Ile	Asn 520	His	Ser	Leu	Ile	Ser 525	Leu	Ser	tta Leu	1584
His	Asp 530	Ile	Ala	Asp	Gln	Leu 535	Tyr	Vaļ	Asp	Ser	Arg 540	Leu	Ser	Val	tat Tyr	1632
Ile 545	Asn	Asp	Ile	Gly	His 550	Lys	Tyr	Cys	Glu	Gln 555	Ile	Ser	Gln	Pro	gga Gly 560	1680
acc Thr	gat Asp	gga Gly	cca Pro	aat Asn 565	act Thr	gaa Glu	gcg Ala	tct Ser	aat Asn 570	Gly	gga Gly	gca Ala	gca Ala	Pro	atc Ile	1728
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<212> PRT

<213> Macaca mulatta rhadinovirus 17577

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520 His Asp Ile Ala Asp Gln Leu Tyr Val Asp Ser Arg Leu Ser Val Tyr 540 535 Ile Asn Asp Ile Gly His Lys Tyr Cys Glu Gln Ile Ser Gln Pro Gly 555 550 Thr Asp Gly Pro Asn Thr Glu Ala Ser Asn Gly Gly Ala Ala Pro Ile 565 570 <210> 84 <211> 2373 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <221> CDS <222> (1)..(2373) <400> 84 atg gag agt tee gte gga tgg ace aaa eae gte gaa eea aat eeg ggg Met Glu Ser Ser Val Gly Trp Thr Lys His Val Glu Pro Asn Pro Gly ttc atc ttg aac atg acg tcc gat gcc aaa gtc agg ggt gtc gtg gat 96 Phe Ile Leu Asn Met Thr Ser Asp Ala Lys Val Arg Gly Val Val Asp 25 cac gtc agt cgc ctg tca aat ata act acc agc cca ccg gaa atg ggt His Val Ser Arg Leu Ser Asn Ile Thr Thr Ser Pro Pro Glu Met Gly tgg tac gac ctg gcc ttc gat ccg gct gaa gac tcc ggg ccg ttc ttg 192 Trp Tyr Asp Leu Ala Phe Asp Pro Ala Glu Asp Ser Gly Pro Phe Leu 50 55 ccq ttt acc gtt tat cta att acg gga act gct ggt gct ggg aaa agt Pro Phe Thr Val Tyr Leu Ile Thr Gly Thr Ala Gly Ala Gly Lys Ser 75 65 acc agc ata tog goo otg tac caa aat tta aac tgo otg atc acg ggo 288 Thr Ser Ile Ser Ala Leu Tyr Gln Asn Leu Asn Cys Leu Ile Thr Gly 90 85 gcg acc acc ata gcc gca cag aac cta tcg cgt cgc cta aag acg ttc 336 Ala Thr Thr Ile Ala Ala Gln Asn Leu Ser Arg Arg Leu Lys Thr Phe 105 100 tgt ccc acg atc ttc agc gct ttt ggc ttt aag agc cga cac atc aat 384 Cys Pro Thr Ile Phe Ser Ala Phe Gly Phe Lys Ser Arg His Ile Asn 120 125 ata gcc gtc aga aaa gct cat cag acc gga gcc gta tcc ata gag caa 432 Ile Ala Val Arg Lys Ala His Gln Thr Gly Ala Val Ser Ile Glu Gln 135 att cag caa cag gag cta tcg aag tat tgg ccg gtt ata gtg gac att 480 Ile Gln Gln Glu Leu Ser Lys Tyr Trp Pro Val Ile Val Asp Ile

505

525

Leu Thr His Met Asp Leu Ile Asn His Ser Leu Ile Ser Leu Ser Leu

500

515

150

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tcc Ser	aac Asn	gcg Ala	aat Asn 180	ttt Phe	gaa Glu	acc Thr	ctc Leu	tcg Ser 185	aga Arg	atg Met	acc Thr	gga Gly	ccg Pro 190	tgt Cys	tta Leu	576
tgg Trp	act Thr	tcc Ser 195	aat Asn	att Ile	att Ile	gta Val	atc Ile 200	gac Asp	gag Glu	gcc Ala	gga Gly	acc Thr 205	ctg Leu	tcc Ser	tct Ser	624
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cta Leu 225	aat Asn	acc Thr	cct Pro	ctt Leu	tac Tyr 230	cgc Arg	cag Gln	999 Gly	gcg Ala	gtt Val 235	ccg Pro	tgc Cys	ata Ile	gta Val	tgc Cys 240	720
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cct Pro	tta Leu	gaa Glu	tac Tyr 340	att Ile	ggc Gly	tgg Trp	acc Thr	cga Arg 345	ctc Leu	ttt Phe	ttg Leu	tca Ser	cat His 350	Ser	gag Glu	1056
gta Val	aag Lys	gcg Ala 355	tat Tyr	cta Leu	aca Thr	aac Asn	cta Leu 360	His	aca Thr	tgt Cys	cta Leu	acg Thr 365	Leu	ggg Gly	Gly ggc	1104
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Glu 545	Leu	Asn	Leu	gaa Glu	Asp 550	Asp	Ile	Phe	Tyr	His 555	Val	Cys	Ser	Pro	Pro 560	1680
cca Pro	ccc Pro	gcg Ala	ggt Gly	atc Ile 565	acc Thr	tcc Ser	ctc Leu	cag Gln	gtt Val 570	ttg Leu	gtc Val	gac Asp	acg Thr	tac Tyr 575		1728
gcc Ala	cta Leu	aag Lys	gac Asp 580	gtg Val	ttc Phe	gcc Ala	tcc Ser	aga Arg 585	Ile	aag Lys	gtg Val	gcg Ala	tgt Cys 590	Arg	tgg Trp	1776
ttt Phe	ggc Gly	ggg Gly 595	gag Glu	ttt Phe	gag Glu	aag Lys	gaa Glu 600	acg Thr	ttt Phe	tcc Ser	gcg	Phe 605	Thr	gtt Val	aac Asn	1824
atg Met	gtc Val 610	Val	agg Arg	gac Asp	gga Gly	gtt Val 615	gac Asp	ttt Phe	gtc Val	tcc Ser	e cct Pro 620	Ser	gaa Glu	cgt Arg	ctc Leu	1872
aac Asn 625	Gly	ctg Leu	ttg Leu	gcg Ala	ttt Phe 630	Ala	tcg Ser	acc Thr	gtt	gaa Glu 635	Ser	tat Tyr	aaa Lys	att Ile	aag Lys 640	1920
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Gly	Tyr	Thr	Phe	Leu 645	Pro	Val	Ala	Phe	Gly 650	Arg	Cys	Gln	Gly	Leu 655	Pro	
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Leu	Ser	Asp	Asp 660	Leu	Arg	Lys	Lys	Met 665	Pro	Ser	Leu	Val	Val 670	Gln	Asp	
tct	agc	ggt Gly	ttt Phe	atc Tle	gcg Ala	tgc Cvs	cta Leu	gag Glu	aat Asn	aac Asn	ata Ile	acc Thr	aaa Lvs	ttg Leu	acc Thr	2064
261	Ser	675	1110	110	ALG	CYB	680	oru.	11011			685	-1 -			
паа	acc	atg	aaa	gac	aaa	agc	att	ttc	caa	ata	tqc	tat	qcq	aaa	qac	2112
Glu	Thr	Met	Glu	Asp	Gly	Ser	Val	Phe	Gln	Val	Cys	Cys	Ala	Gly	Asp	
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tat	a gg	gtc	agc	tca	aat	tta	gcc	atg	acc	atc	gta	aag	gca	cag	gga	2160
Tyr 705	Gly	Val	Ser	Ser	Asņ 710	Leu	Ala	Met	Thr	715	vaı	гÃг	Ата	GIII	720	
										~~~	+ 00	<b>6</b> 2.6	220	220	ata	2208
atg Met	Ser	ttg Leu	gag Glu	Arg	gca Val	Ala	Val	Val	Phe	Gly	Ser	His	Lys	Asn	Val	2200
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Gln	Thr	Ser	His 740	Val	Tyr	Val	Ala	Ile 745	Ser	Arg	Ala	Val	Asn 750	Ser	Asn	
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tat	ttg	gtc Val	atg Met	gac	agc Ser	aac Asn	ccc Pro	ctt Leu	aaa Lvs	acc Thr	ctc Leu	ctc Leu	aga Arq	gaa Glu	cca Pro	2304
1 7 1	ыса	755		1100	001		760	204	_, _			765				
at.c	αat	aac	acc	tcc	qcc	aaq	cat	ata	qtc	cqc	gcc	ctc	cac	aac	сса	2352
Val	Asp	Asn	Thr	Ser	Āla	Lys	His	Ile	Val	Arg	Ala	Leu	His	Asn	Pro	
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His	Val	Ser 35	Arg	Leu	Ser	Asn	.Ile 40	Thr	Thr	ser	Pro	Pro 45	Glu	Met	GIY	
Trp		Asp	Leu	Ala	Phe		Pro	Ala	Glu	Asp		Gly	Pro	Phe	Leu	
Pro	50 Phe	Thr	Val	Tyr	Leu	55 Ile	Thr	Gly	Thr	Ala	60 Gly	Ala	Gly	Lys	Ser	
65					70					75					80	
Thr	Ser	Ile	ser	Ala 85	ьeu	Tyr	GIN	Asn	Leu 90	Asn	cys	ьeu	тте	Thr 95	GTÀ	
Ala	Thr	Thr			Ala	Gln	Asn		Ser	Arg	Arg	Leu		Thr	Phe	
Cvs	Pro	Thr	100 Ile	Phe	Ser	Ala	Phe	105 Gly	Phe	Lys	Ser	Arg	110 His	Ile	Asn	
- 1	. –							4		-						

		115					120					125			
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145	Gln		Gln		150					155					160
			Val	165					170					175	
			Asn 180					185					190		
_		195	Asn				200					205			
-	210		Thr			215					220				
225			Pro		230					235					240
	_		Pro	245					250					255	
_			Lys 260					265					270		
		275	Lys				280					285			
	290		Phe			295					300				
305			Lys		310					315					320
			Val	325					330					335	
			Tyr 340					345					350		
		355	Tyr				360					365			
_	370		Asp			375					380				
385			Pro		390					395					400
			Thr	405					410					415	
_			Phe 420					425					430		
		435	Ser				440					445			
	450		Ser			455					460				
465			Tyr		470					475					480
			His	485					490					495	
			500					505					510		Val
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	530					535					540				Glu
Glu 545	Leu	Asn	Leu	Glu	Asp 550	Asp	Ile	Phe	Tyr	His 555	Val	Cys	Ser	Pro	Pro 560
Pro			Gly	565					570					575	
Ala	Leu	Lys	Asp 580		Phe	Ala	Ser	Arg 585	Ile	Lys	Val	Ala	Cys 590		Trp
Phe	Gly	Gly 595	Glu	Phe	Glu	Lys	Glu 600	Thr	Phe	Ser	Ala	Phe 605		Val	Asn

Met Val Val Arg Asp Gly Val Asp Phe Val Ser Pro Ser Glu Arg Leu 615 620 Asn Gly Leu Leu Ala Phe Ala Ser Thr Val Glu Ser Tyr Lys Ile Lys 630 635 Gly Tyr Thr Phe Leu Pro Val Ala Phe Gly Arg Cys Gln Gly Leu Pro 650 645 Leu Ser Asp Asp Leu Arg Lys Lys Met Pro Ser Leu Val Val Gln Asp 660 665 Ser Ser Gly Phe Ile Ala Cys Leu Glu Asn Asn Ile Thr Lys Leu Thr 680 685 Glu Thr Met Glu Asp Gly Ser Val Phe Gln Val Cys Cys Ala Gly Asp 700 695 Tyr Gly Val Ser Ser Asn Leu Ala Met Thr Ile Val Lys Ala Gln Gly 715 710 Met Ser Leu Glu Arg Val Ala Val Val Phe Gly Ser His Lys Asn Val 730 725 Gln Thr Ser His Val Tyr Val Ala Ile Ser Arg Ala Val Asn Ser Asn 740 745 Tyr Leu Val Met Asp Ser Asn Pro Leu Lys Thr Leu Leu Arg Glu Pro 760 Val Asp Asn Thr Ser Ala Lys His Ile Val Arg Ala Leu His Asn Pro 775 Asn Thr Thr Leu Ile Tyr

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-						_							gac Asp 110			336
			-	_						-			tcg Ser			384
_										-		_	ttt Phe			432
_			_			_					_	-	ccc Pro	_		480
													tac Tyr			528
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Pro	Ala	Trp 195	Ser	Gly	Asp	Ile	Ser 200	Arg	Ser	Pro	Ala	Glu 205	ggc	Gly	Trp	624
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	-		_				_		_	_	_		cgc Arg	-		864
													ggt Gly			912
													G] y			960
													atg Met			1008
acc	ccc	ccg	gtg	tgt	gga	aat	gac	aac	tat	ccg	tgg	ccg	tgg	ttg	gac	1056

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tga 1059

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ctt gaa ttt tt Leu Glu Phe Le 35	a aat ctt tc u Asn Leu Se	t cca ttt t r Pro Phe L 40	ta aaa cag aag o eu Lys Gln Lys I 45	ctc gcg gct 144 Leu Ala Ala
ctg ctg aag cg Leu Leu Lys Ar 50	g Val Met As	t atg agc a p Met Ser A 5	ac gta acc gtg a sn Val Thr Val 1	att tat cca 192 Ile Tyr Pro
ccg ata gat ag Pro Ile Asp Ar 65	a att atg tg g Ile Met Tr 70	g tgg tcg t p Trp Ser T	at tgt tgc gaa yr Cys Cys Glu 75	ccg gag gat 240 Pro Glu Asp 80
att aaa gtc gt Ile Lys Val Va	g atc ctt gg l Ile Leu Gl 85	y Gln Asp P	ct tac cat cgc ro Tyr His Arg 90	ggt caa gcc 288 Gly Gln Ala 95
acc gga cta gc Thr Gly Leu Al 10	a Phe Ser Va	t gct ccg g l Ala Pro A 105	ac tac agt ata .sp Tyr Ser Ile	cct cca agc 336 Pro Pro Ser 110
ctc aaa aat at Leu Lys Asn Il 115	t ttt aaa ga e Phe Lys Gl	g ata gcc a u Ile Ala A 120	at act gta cct sn Thr Val Pro 125	ggg ttc acc 384 Gly Phe Thr
gct cct tct ca Ala Pro Ser Hi 130	c ggg tgc tt s Gly Cys Le 13	u Asp Cys T	gg gca aaa cgg rp Ala Lys Arg 140	gga gtt ctg 432 Gly Val Leu
ctt tta aac ac Leu Leu Asn Th 145	c att ctg ac r Ile Leu Th 150	g gtg gaa a r Val Glu A	ga ggg aag gcg arg Gly Lys Ala 155	ggg tca cac 480 Gly Ser His 160
gcc aac ctt gg Ala Asn Leu Gl	c tgg gat tg y Trp Asp Tr 165	p Phe Thr S	gc tac ata ata er Tyr Ile Ile 70	agc tgc ctt 528 Ser Cys Leu 175
tct gcc aag ct Ser Ala Lys Le 18	u Gln Arg Cy	c gtt ttt a s Val Phe M 185	itg ctg tgg gga Met Leu Trp Gly	aga aag gct 576 Arg Lys Ala 190
ata gac aag gc Ile Asp Lys Al 195	g acg ctg at a Thr Leu Il	a aac gga c e Asn Gly G 200	ag aga cat ctċ In Arg His Leu 205	gtc ctc aag 624 Val Leu Lys

672

720

768

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Ala Arg His Pro Ser Pro Leu Ala Thr Ala His Ala Ala Thr Gly Ser
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Pro Trp Pro Gln Phe Leu Gly Cys Asn His Phe Lys Leu Ala Asn Asp
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Val Leu Glu Glu Pro Ser Thr Gln Thr Leu Leu Leu Ser Asp Ser Trp
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Leu Glu Phe Leu Asn Leu Ser Pro Phe Leu Lys Gln Lys Leu Ala Ala
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                           40
Leu Leu Lys Arg Val Met Asp Met Ser Asn Val Thr Val Ile Tyr Pro
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Pro Ile Asp Arg Ile Met Trp Trp Ser Tyr Cys Cys Glu Pro Glu Asp
Ile Lys Val Val Ile Leu Gly Gln Asp Pro Tyr His Arg Gly Gln Ala
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Thr Gly Leu Ala Phe Ser Val Ala Pro Asp Tyr Ser Ile Pro Pro Ser
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Leu Lys Asn Ile Phe Lys Glu Ile Ala Asn Thr Val Pro Gly Phe Thr
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Ala Pro Ser His Gly Cys Leu Asp Cys Trp Ala Lys Arg Gly Val Leu
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Leu Leu Asn Thr Ile Leu Thr Val Glu Arg Gly Lys Ala Gly Ser His
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Ala Asn Leu Gly Trp Asp Trp Phe Thr Ser Tyr Ile Ile Ser Cys Leu
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Ser Ala Lys Leu Gln Arg Cys Val Phe Met Leu Trp Gly Arg Lys Ala
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Ile Asp Lys Ala Thr Leu Ile Asn Gly Gln Arg His Leu Val Leu Lys
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                           200
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Ala Arg His Pro Ser Pro Leu Ala Thr Ala His Ala Ala Thr Gly Ser
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Pro Trp Pro Gln Phe Leu Gly Cys Asn His Phe Lys Leu Ala Asn Asp
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ata Ile	cca Pro	ctt Leu 35	agt Ser	gac Asp	ttt Phe	ata Ile	ttt Phe 40	ccg Pro	gag Glu	ccg Pro	ttt Phe	gag Glu 45	att Ile	gct Ala	tct Ser	144
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ctt Leu	gat Asp	gag Glu 115	ttt Phe	tta Leu	gcc Ala	gag Glu	ttt Phe 120	gaa Glu	gac Asp	ttt Phe	cac His	ata Ile 125	aac Asn	ggt Gly	agt Ser	384
gaa Glu	agc Ser 130	gga Gly	act Thr	gct Ala	tat Tyr	acg Thr 135	cgg Arg	cca Pro	cct Pro	ctg Leu	ttg Leu 140	gat Asp	ttc Phe	tca Ser	gac Asp	432
aga Arg 145	agt Ser	aca Thr	aaa Lys	gtt Val	tca Ser 150	cgt Arg	ata Ile	cgt Arg	aaa Lys	gta Val 155	att Ile	acc Thr	aga Arg	cgc Arg	999 Gly 160	480
		tgg Trp							tag 170							510

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<211> 169

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<213> Macaca mulatta rhadinovirus 17577

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gaa tot oot atg goo gag ata tgg ogo gao tgo aaa gag ogo tto tgo 336 Glu Ser Pro Met Ala Glu Ile Trp Arg Asp Cys Lys Glu Arg Phe Cys 105 ctc gct ctg gaa ctg gcg tgc ggc tgt caa agg tgc gcg agc gcc gcc 384 Leu Ala Leu Glu Leu Ala Cys Gly Cys Gln Arg Cys Ala Ser Ala Ala 115 120 432 agg cag cta cgg gcc tgt cag caa gcc tgc agg cca cct aag ctg aat

Arg Phe Val Thr Gln Gln Ile Tyr Met His Leu Lys Asp His Ala Ser

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						act Thr										528
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gtc Val	cgg Arg	gtg Val 195	gaa Glu	acc Thr	agc Ser	ege Arg	gtt Val 200	gcc Ala	tcc Ser	tgt Cys	ttg Leu	aac Asn 205	ttg Leu	tcg Ser	tgg Trp	624
tta Leu	tac Tyr 210	ttg Leu	att Ile	tta Leu	gac Asp	tcg Ser 215	tat Tyr	gtt Val	cga Arg	aca Thr	gat Asp 220	tta Leu	aca Thr	aat Asn	ctg Leu	672
gaa Glu 225	atg Met	gca Ala	atg Met	agc Ser	cgt Arg 230	gcc Ala	tgc Cys	cgc Arg	att Ile	cac His 235	ggc Gly	ctt Leu	agc Ser	gcc Ala	999 Gly 240	720
gac Asp	ccg Pro	ttt Phe	tat Tyr	tcc Ser 245	gcc Ala	ctc Leu	gtg Val	tgg Trp	tta Leu 250	aaa Lys	aat Asn	agt Ser	tac Tyr	gca Ala 255	tgt Cys	768
gac Asp	acg Thr	aat Asn	aca Thr 260	ttt Phe	ttt Phe	ttc Phe	acc Thr	gtc Val 265	aat Asn	tca Ser	acc Thr	agt Ser	gtc Val 270	acg Thr	act Thr	816
						tgt Cys										864
						ctg Leu 295										912
						gcc Ala										960
						gat Asp										1008
						ctg Leu										1056
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Phe Glu Glu Leu Ser 385

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	> CI	os L)	(906)	,	•											
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acc Thr	gtc Val	aat Asn 35	gcc Ala	cgc Arg	aat Asn	ccg Pro	ctc Leu 40	tac Tyr	caa Gln	gcg Ala	gca Ala	acc Thr 45	ctt Leu	aga Arg	gtg Val	144
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agc Ser 145	tgg Trp	tgc Cys	cta Leu	tct Ser	cac His 150	atg Met	gtc Val	ggt Gly	gta Val	acc Thr 155	aaa Lys	acc Thr	ttc Phe	aaa Lys	999 Gly 160	480
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gtt ata tac gag gac tac cag gac acg cag ttt aac gtg ttt tta aat Val Ile Tyr Glu Asp Tyr Gln Asp Thr Gln Phe Asn Val Phe Leu Asn ctt tgt ttt ttt tgg acc act gtc ata aag atg tac cag agt tgc att Leu Cys Phe Phe Trp Thr Thr Val Ile Lys Met Tyr Gln Ser Cys Ile ttt aaa gac aag cta ttg gac acg att aaa gct tgc ata gag ctt cta Phe Lys Asp Lys Leu Leu Asp Thr Ile Lys Ala Cys Ile Glu Leu Leu aaa ggc gag gcc agg cag ttt ttt ggt tgg tac gac cta aac acg cca Lys Gly Glu Ala Arg Gln Phe Phe Gly Trp Tyr Asp Leu Asn Thr Pro aat tta ggt tca tcg gca cta gta aag tac aca gag cac ctg atc cga Asn Leu Gly Ser Ser Ala Leu Val Lys Tyr Thr Glu His Leu Ile Arg qua eta agt gtg gat tea tea gee att eee att gge gag ata tge tee Ala Leu Ser Val Asp Ser Ser Ala Ile Pro Ile Gly Glu Ile Cys Ser cac cta cac cac tgt aaa cac gcc ctc ctg aat ctt gaa taa His Leu His His Cys Lys His Ala Leu Leu Asn Leu Glu <210> 95 <211> 301 <212> PRT <213> Macaca mulatta rhadinovirus 17577 Met Ser Arg His Tyr Gly Lys Asp His Leu Leu Asn His Met Tyr Lys Phe His Tyr Pro Pro Leu Gly Met Ile Val Gly Glu Met Asn Thr Leu Thr Val Asn Ala Arg Asn Pro Leu Tyr Gln Ala Ala Thr Leu Arg Val Glu Arg Ala Leu Tyr Leu Ser Lys Ile Leu Gln Val Leu Met Gln His Arg Gln Gly Glu Arg Phe Ile Val Pro Gln Cys Arg Ser Asn Met Val Tyr Cys Leu Lys Glu Leu His Lys Ile Thr Asn Asp Arg Ile Arg Gly Leu Ile Asn Ser Val Leu Pro Leu Val Asp Ala Gly Cys Val Gly Phe Asp Glu Glu Leu Val Arg Val Leu Pro Glu Ile Leu Lys Leu Glu Tyr Pro His Ala His Glu Leu Leu Pro Pro His Asp Pro Thr Ser Pro Leu

Ser Trp Cys Leu Ser His Met Val Gly Val Thr Lys Thr Phe Lys Gly

Glu Val Lys Glu Met Ile Asp Thr Phe His Asp Leu Ser Val Pro Ser

Phe Gln Tyr Leu Ala Ser Leu Val Lys Lys Phe Phe Leu Val Glu Glu

Val Ile Tyr Glu Asp Tyr Gln Asp Thr Gln Phe Asn Val Phe Leu Asn 200 205 195 Leu Cys Phe Phe Trp Thr Thr Val Ile Lys Met Tyr Gln Ser Cys Ile 215 210 Phe Lys Asp Lys Leu Leu Asp Thr Ile Lys Ala Cys Ile Glu Leu Leu 235 230 Lys Gly Glu Ala Arg Gln Phe Phe Gly Trp Tyr Asp Leu Asn Thr Pro 245 250 Asn Leu Gly Ser Ser Ala Leu Val Lys Tyr Thr Glu His Leu Ile Arg 270 265 260 Ala Leu Ser Val Asp Ser Ser Ala Ile Pro Ile Gly Glu Ile Cys Ser 280 285 His Leu His His Cys Lys His Ala Leu Leu Asn Leu Glu 295

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			gat Asp									624
			ctg Leu									672
		_	gcc Ala			_	_				_	720
	_	_	cct Pro	_		_	_					768
			caa Gln 260									816
			cag Gln									864
			gcg Ala	_					 _			 912
			cgg Arg									960
	_	_	gtt Val	_								1008
			acg Thr 340									1056
			ggc Gly									1104
			tcc Ser									1152

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	Asp	Ser		100 Leu	Val	Thr	Tyr		105 Thr	Leu	Asp	Ala		110 Gly		Leu	
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Ile Leu Ser Ala Pro Pro Leu Ser Gln Phe Val Ile Thr Asn Thr His
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Pro Ser Leu Pro Gln Ser Val Ser Ile Ile Thr Pro Thr Gln Gly Val
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Val Pro Gly Gln Cys Phe Met Asp Thr Trp Lys Ala Val Ser Gln Ser
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Ile His His Gln Ala Gln Thr Pro Ile Leu Ala Ala Ala Leu Thr Gly
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Ser Thr Ser Ala Ala Pro Gly Pro His Ile Ala Cys Ser Pro Val Ala
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Pro Ala Cys Val Pro Gln Pro Ala Leu Pro Pro Asn Val Pro Ala Lys
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Arg Met Glu Thr Val Ala Gln Leu Gly Asn Ala Pro Val Lys Asn Val
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His Ile Gly Gly Arg Val Tyr Ala Pro Leu Val Asn Ile Pro Ile Ile
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Asp Leu Thr Ser Pro Ser Gly Ser Gly Gln Ser Pro Ala Asp Ile Ala
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Asn Thr Pro Glu Ser Arg Met Ala Ala Gly Ser Pro Pro Phe Ala Glu
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Thr Ala Ala Thr Val Pro Ala Lys Arg Lys Gln Pro Arg Glu Asp Val
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Ala Asp Lys Arg Leu Lys Gly Asp Val Arg Gly Ala Ala Thr Val Asn
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His Pro Phe Pro Gly Pro Ser Gly Met Arg Val Arg Glu Gln Gly Leu
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Phe Asp Leu Ile Glu Ser Ser Thr Asp Val Thr Ala Asn Ala Ser Gly
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                                      460
Pro Lys Asn Asp Asp Met Leu Ala Ala Ile Leu Gln Asp Leu Tyr
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Gly Leu Gln Ser Pro Pro Ala Ile Asp Ser Pro Ser Ser Asn Ser Asp
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gag Glu	aat Asn 50	aac Asn	ggc Gly	ccg Pro	ttt Phe	tcc Ser 55	caa Gln	ata Ile	atg Met	cac His	aat Asn 60	gga Gly	cag Gln	agc Ser	aat Asn	192
acc Thr 65	ggg Gly	aca Thr	ggt Gly	gaa Glu	agc Ser 70	ttc Phe	ggc Gly	agc Ser	tac Tyr	gct Ala 75	gcc Ala	ggc Gly	gac Asp	ggt Gly	ttt Phe 80	240
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Gly aaa	gca Ala 130	cat His	gcc Ala	gta Val	tct Ser	gac Asp 135	cgg Arg	ata Ile	ggc Gly	aga Arg	gac Asp 140	ggt Gly	ggc Gly	gct Ala	gac Asp	432
aat Asn 145	aga Arg	cta Leu	ctc Leu	aag Lys	gtg Val 150	agt Ser	gcg Ala	cgg Arg	ctg Leu	tcg Ser 155	gac Asp	aaa Lys	aca Thr	aag Lys	agc Ser 160	480
gcc Ala	ctt Leu	cgc Arg	agc Ser	cat His 165	cct Pro	tgc Cys	ttg Leu	cgt Arg	tgc Cys 170	tat Tyr	tct Ser	ttg Leu	atg Met	ttt Phe 175	aac Asn	528
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1		Leu		5				Thr	10				Asp	15 Ser		
Asn	Leu	Cys	20 Pro	Asp	Gly	Gln	_	25 Leu	Leu	Gly	Ser				Thr	
Glu	Asn	35 Asn	Gly	Pro	Phe	Ser	40 Gln	Ile	Met	His	Asn	45 Gly		Ser	Asn	

Thr Gly Thr Gly Glu Ser Phe Gly Ser Tyr Ala Ala Gly Asp Gly Phe 70 75 Leu Gly Gly Ser Val Ser Gly Met Tyr Gly Asn Asn Thr Gly Glu Gly 90 Ala Cys Ser Lys Arg Pro Ser Ala Cys Arg Lys Arg Ser Ala Ala Leu 105 100 Ile His Ala Ala Ser Glu Ala Ser Val Ala Glu Gln Gly Thr Ser Gln 125 115 120 Gly Ala His Ala Val Ser Asp Arg Ile Gly Arg Asp Gly Gly Ala Asp 140 135 Asn Arg Leu Leu Lys Val Ser Ala Arg Leu Ser Asp Lys Thr Lys Ser 155 . 150 Ala Leu Arg Ser His Pro Cys Leu Arg Cys Tyr Ser Leu Met Phe Asn 1.65 170 Thr <210> 100 <211> 693 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <221> CDS <222> (1)..(693) <400> 100 atg gga ttt ggg aac ata cgt ctg gga tgg agg tta tgc ttc atg gtc Met Gly Phe Gly Asn Ile Arg Leu Gly Trp Arg Leu Cys Phe Met Val tgg gtg gcg tgg att gca cgg gga cgg tcg gtg tgc cca acc tgg cac 96 Trp Val Ala Trp Ile Ala Arg Gly Arg Ser Val Cys Pro Thr Trp His 25 ctg aca gat ggg aaa tac gag gcg gta tac agg cac tac ctc gaa gag Leu Thr Asp Gly Lys Tyr Glu Ala Val Tyr Arg His Tyr Leu Glu Glu 40 192 tgc cgc aaa cat gaa ggc tcg ggg agc ctg gac ggt tcc gga cag aca Cys Arg Lys His Glu Gly Ser Gly Ser Leu Asp Gly Ser Gly Gln Thr aag ggg tot gga acc aaa gca acc acc gaa got aat ata tog ata aga Lys Gly Ser Gly Thr Lys Ala Thr Thr Glu Ala Asn Ile Ser Ile Arg 70 cet aac git gic aca ica ggi caa aat aaa gag eeg eei ggg aca gea 288 Pro Asn Val Val Thr Ser Gly Gln Asn Lys Glu Pro Pro Gly Thr Ala 85 ccq agg gcc gaa tca tca cac gac ctg cca egc atc aag cag gtt aac 336 Pro Arg Ala Glu Ser Ser His Asp Leu Pro Arg Ile Lys Gln Val Asn 105 get etc ega tta tea acc eeg gaa ttg geg caa eea etc eeg gta gta Ala Leu Arg Leu Ser Thr Pro Glu Leu Ala Gln Pro Leu Pro Val Val

55

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gtg Val	gtc Val	ctg Leu	tct Ser	ttc Phe 165	aga Arg	gcc Ala	atc Ile	cgt Arg	gcg Ala 170	cgg Arg	tcc Ser	aca Thr	cgc Arg	gat Asp 175	acc Thr	528
gag Glu	cag Gln	tcc Ser	gtt Val 180	cgc Arg	gat Asp	cgg Arg	aac Asn	acg Thr 185	gtc Val	acg Thr	acc Thr	agc Ser	tat Tyr 190	cgt Arg	acc Thr	576
cct Pro	ggc Gly	cgc Arg 195	cct Pro	tcc Ser	ctc Leu	ttt Phe	caa Gln 200	gcc Ala	aga Arg	ccc Pro	tcg Ser	tct Ser 205	cac His	ggt Gly	gcg Ala	624
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	ata Ile			_		tga										693
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Arg Leu Pro Pro Ser Pro Arg Thr Met Ala Arg Tyr Ala Glu Ser Arg

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Met Tyr Ala Ser Pro Gly Pro Leu Lys Cys Pro Ile Leu Asn Leu Pro

		115					120					125				
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gat Asp	ata Ile 210	cca Pro	gca Ala	aac Asn	agc Ser	cgc Arg 215	ata Ile	tgt Cys	cag Gln	gta Val	gtg Val 220	ttt Phe	atc Ile	cac His	gaa Glu	672
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gcc Ala	acg Thr	Leu	ccc Pro 260	aaa Lys	acc Thr	cac His	ccg Pro	ctc Leu 265	aac Asn	tcc Ser	cgc Arg	cac His	act Thr 270	caa Gln	agc Ser	816
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	tta Leu 290	taa														873
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Asn	Lys			Val	Val	Val	Lys 40			Glu	Pro	Leu 45	ı Val		Pro	
Leu	Gly 50		Lys	Ile	Ile	Arg 55	Ala	Pro	Gln	Сув	Ala 60	Phe		e Let	ı Ser	

Gly Ala Pro Thr Asp Glu Val Tyr Tyr His Thr Gly Leu Ile Asp Gln 70 75 Gly Tyr Arg Gly Glu Ile Lys Leu Ile Val Leu Asn Lys Thr Lys Gln 85 90 Val Val Thr Leu Tyr Arg Gly Glu Val Asn Val Ser Leu Ile Ala Phe 105 Met Tyr Ala Ser Pro Gly Pro Leu Lys Cys Pro Ile Leu Asn Leu Pro 115 120 125 His Tyr Ser Leu Asp Ala Gly Phe Asp Val Thr Ser Pro His Ala Met 135 140 Thr Ile Pro Pro Thr Asp Arg Thr Pro Phe Thr Leu Ser Leu Tyr Tyr 155 Lys Ser Pro Gln Leu Ser Thr Pro His Val Pro Leu Ile Val Gly Arg 170 175 165 Ser Gly Leu Ala Thr Lys Gly Leu Thr Val Asp Ala Thr Lys Trp Thr 190 180 185 Gln Ser Leu Val His Leu Arg Phe Tyr Asn Phe Thr Lys Glu Pro Ile 200 205 Asp Ile Pro Ala Asn Ser Arg Ile Cys Gln Val Val Phe Ile His Glu 215 220 Asp His Val Pro Ser Gly Trp Asn Ile Leu Arg Ser Arg Val Gln Leu 235 225 230 Gly Ser Thr Leu Gln Ile Ser Trp Ala Lys Ile Arg Phe Thr Asp Val 245 250 255 Ala Thr Leu Pro Lys Thr His Pro Leu Asn Ser Arg His Thr Gln Ser 265 270 Gln Thr Glu Pro Glu Thr Ala Arg Gly Ala Lys Gly Leu Gly Ser Ser 280 Gly Leu 290 <210> 108 <211> 633 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(633)

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	gaa aaa ctg Glu Lys Leu 150				
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Val Asp Met 50	Gly Asp Val	Lys Gln Ala 55	Glu Met Cys 60	Thr Ala Ala	Leu
65	Tyr Leu Leu .		75		80
Leu Arg Arg	Phe Asp Ala 85	Ala Arg Val	Pro Ala Gly 90	Cys Gln Glu 95	
	Gln Ile Ser 100	105		110	
Asn Ala Met 115	Leu Ser Leu	Ala Ile Gly 120	Asp Ile Thr	Val Asp Glu 125	Ser

135 Leu Glu Met Glu Lys Leu Ala Thr Thr Ile Ala Ser Asp Asp Ser Val 150 155 160 Thr Trp Ala Ala Glu Ile Asn Asn Val Leu Val Asp Thr Glu Ala Ser 170 165 Ser Asn Pro Ser His Pro Val Ile Arg Gln Pro Thr Pro Gln Leu Ala 1.90 185 Val Ala Asp Asn Ile Val Pro Asp Lys Ile Ile Gln Asp Ala Gln Ala 205 195 Asp Gly 210 <210> 110 <211> 2487 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(2487) <400> 110 atg gta gat gaa att agg gca att ttc tct act agt gga gat atg gcc Met Val Asp Glu Ile Arg Ala Ile Phe Ser Thr Ser Gly Asp Met Ala 10 96 gaa gta att acg gat ata ctg act gaa acg caa gca acg gcg tcc ttc Glu Val Ile Thr Asp Ile Leu Thr Glu Thr Gln Ala Thr Ala Ser Phe tto tgo gtg oto cao gat ogg ggo gao gog oot ata aat act ooa oat Phe Cys Val Leu His Asp Arg Gly Asp Ala Pro Ile Asn Thr Pro His 40 que qua att aaa etc tge etg eec gee aag ege eea gge gge ggg eea 192 Ala Val Ile Lys Leu Cys Leu Pro Ala Lys Arg Pro Gly Gly Pro 55 agg tgt tta ccg ttg atg gtg ctg aac cta ccg gcg tgg cag gtt aat Arg Cys Leu Pro Leu Met Val Leu Asn Leu Pro Ala Trp Gln Val Asn 70 65 cta ttc cta aca ggt gac gca cca ttg acc tcg gat aac att aaa gac 288 Leu Phe Leu Thr Gly Asp Ala Pro Leu Thr Ser Asp Asn Ile Lys Asp 90 85 cgc att gac ctg gct cag acc gag gaa ata ctc gaa ccc ata tta agc 336

Ala Phe His Ala Leu Leu Asn Lys Arg Ala Asp Glu Thr Val Ser Leu

140

432

Arg Ile Asp Leu Ala Gln Thr Glu Glu Ile Leu Glu Pro Ile Leu Ser

gta ctg gca tgc aaa cgg tee geg cag cag acc aaa cat gac teg ttt Val Leu Ala Cys Lys Arg Ser Ala Gln Gln Thr Lys His Asp Ser Phe 120

135

105

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ttc Phe	agc Ser	tcc Ser 195	agt Ser	cag Gln	ggc Gly	cag Gln	agc Ser 200	ttg Leu	gtc Val	acc Thr	gta Val	aac Asn 205	acc Thr	tat Tyr	gac Asp	624
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		gaa aaa cca cta att Glu Lys Pro Leu Ile 715	
		cac gat acc ata cag His Asp Thr Ile Gln 735	
ttc ttt ccg gat gac Phe Phe Pro Asp Asp 740	cgc atc ggc caa ttt Arg Ile Gly Gln Phe 745	gca tct gtg agc ttc Ala Ser Val Ser Phe 750	atg 2256 Met
		cca caa aaa gga aac Pro Gln Lys Gly Asn 765	
		cac act cag aca gtc His Thr Gln Thr Val 780	
		agc gag gtc acg gtg Ser Glu Val Thr Val 795	
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520 525 His Pro Val Tyr Phe Phe Lys Ser Ala Cys Pro Ala Val Thr Trp Pro 540 535 Asp Asp Ile Ser Asp Thr Ala Phe Cys His Cys Asp Ala Lys Ile Gly 550 555 Met Arg Ile Val Thr Pro Phe Pro Ser Gly Tyr Cys Leu Val Gly Ser 565 570 Ala Pro Leu Val Ser Leu Thr Asp Ile Leu Asn Arg Val Val Lys Leu 585 590 Asp Thr Arg Leu Ala Ser Glu Tyr Pro Gly Ile Leu Glu Asp Lys Gly 600 Pro Phe Asp Ser Gly Ile Tyr Ala Lys Gly Arg Cys Val Arg Val Pro 615 620 His Cys Tyr Lys Val Gly Pro Gly Gly Glu Leu Ser Arg Leu Leu Lys 635 630 Ile Ile Ile Cys His Pro Glu Glu Ser Asp Lys Ser Ala Tyr Leu Lys 650 645 Asn Ala Phe Lys Val Ser Asn Leu Leu His His Ala Pro Gly Asp Ser 670 665 660 Val Thr Lys Asn Gly His Leu Val Tyr Ala Ile Thr Asp Glu Asn Glu 680 685 675 Gly Phe Leu Glu Ser Lys Thr Lys Asn Asn Leu Pro Lys Thr Ile Thr 700 695 Asp Leu Ala Glu Lys Ile Glu Arg Thr Thr Glu Lys Pro Leu Ile Asp 715 710 Trp Ala Ala Thr Ala Val Trp Pro Lys Leu His Asp Thr Ile Gln Arg 730 725 Phe Phe Pro Asp Asp Arg Ile Gly Gln Phe Ala Ser Val Ser Phe Met 750 745 His Ser Gly Asp Asn Ile Ile Gln Val Lys Pro Gln Lys Gly Asn Asn 760 765 755 Phe Phe Cys Ile Asn His Lys His Arg Asn His Thr Gln Thr Val Arg 775 Val Phe Leu Thr Leu His Ser Thr Lys Glu Ser Glu Val Thr Val Thr 795 790 Phe Met Ser Gln Cys Phe Ala Ala Lys Cys Asn His Asn Ser Pro Thr 805 810 Ala His Phe Ser Phe Met Val Pro Ile Thr Gly Thr 820

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Ser Ser Asp Phe Asp Asp Ser Ser Ser Asp Glu Met Asp Asp Leu Ser
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gtg Val 225	gcc Ala	aaa Lys	aaa Lys	gtt Val	agc Ser 230	ctg Leu	gct Ala	aaa Lys	cta Leu	aca Thr 235	agc Ser	cta Leu	tac Tyr	aaa Lys	Pro 240	720
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Arg His Arg Glu Pro Trp His Arg Gly Gly Lys Gly Lys Ala Pro Phe
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                      120
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                                    140
Asp Tyr Arg Gly Lys Ala Ala Leu Thr Arg Ser Ile Lys Glu Ser Ile
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Lys Lys Met His Leu Pro Ser Thr Met Leu Ser Arg Ala His Asp Lys
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            165 170
Lys Val Phe Glu Gly Leu Leu Pro Arg His Leu Gly Gln Cys Phe Gln
       180 185
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Val Cys Leu Pro Ala Pro Pro Pro Leu Gln Pro Glu Val Phe Thr Asp
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Arg Gln Leu Thr Ala Ile Val Lys Ser Gly Gly Arg Arg Asp Ala Leu
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Val Ala Lys Lys Val Ser Leu Ala Lys Leu Thr Ser Leu Tyr Lys Pro
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Leu Leu Thr Phe Val Thr Gly Arg Asn Asn Gln Ala His Trp Leu Ala
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Thr Arg Lys Asn Thr Leu Ala Ser Ala Gly Leu Glu Ala Leu Ala Ala
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Phe Ile Glu Glu Gly Leu Ala Trp Ala Gln Val Cys Val Ser Gln Asn
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Ser Val Cys Thr Trp Phe Ile Ser Lys Ile Arg His Leu His Ile Gln
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Cys Phe Leu Glu Asn Gln Gly Glu Val Ser Leu Val Lys Gln Leu Thr
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Tyr Leu Val Cys Ile Asn Asn Arg Leu Ala Glu Ala Ala Asn Leu Ala
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Gly Glu Val Lys Leu Asn Phe Lys Leu Gly Met Leu Ile Gly Phe Ala
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Leu Thr Leu Pro Ala Leu Leu Ala Glu His Lys Leu Ser Gly Glu Ser
   370 375
Leu Tyr Leu Phe Arg Ser Phe Leu Glu Lys Tyr Arg Pro Gly Asp Val
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Met Gly Leu Leu Asn Ser Ile Val Val Glu His Tyr Thr Lys Cys Arg
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										gaa Glu						384
										gaa Glu						432
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acc Thr	Āla	tcc Ser 195	ggt Gly	Glu	gac Asp	Gly	Arg	Gln	gac Asp	aat Asn	agt Ser	caa Gln 205	Gly	ggc Gly	gcg Ala	624
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cat	tct	<b>99</b> 9	atg	ggc	gtt	cgt	tta	tcc	acg	aga	cca	acg	gat	aaa	aat	816

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	50					55					60				Tyr	
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<222> (1)..(1173)

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<400> 124

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		aac Asn														192
caa Gln 65	acc Thr	ggt Gly	tca Ser	cat His	gcc Ala 70	cag Gln	aaa Lys	ttt Phe	aag Lys	aaa Lys 75	att Ile	agg Arg	atg Met	tta Leu	tat Tyr 80	240
gca Ala	gtg Val	aga Arg	tct Ser	cac His 85	agg Arg	tat Tyr	ttg Leu	agg Arg	gag Glu 90	ctg Leu	aca Thr	ccg Pro	ccg Pro	agc Ser 95	aag Lys	288
gcc Ala	gly ggg	ggc Gly	gtc Val 100	tct Ser	gly ggg	gaa Glu	aga Arg	tac Tyr 105	aga Arg	ctc Leu	ttt Phe	caa Gln	ttg Leu 110	ctt Leu	cct Pro	336
gag Glu	gtt Val	acg Thr 115	gtg Val	ggc Gly	tgc Cys	gat Asp	ctg Leu 120	tgt Cys	aac Asn	ctc Leu	atc Ile	gcg Ala 125	acc Thr	aca Thr	tcg Ser	384
ttg Leu	cat His 130	agc Ser	tgt Cys	tcc Ser	atg Met	ggc Gly 135	agt Ser	tgc Cys	gtt Val	cga Arg	gag Glu 140	gat Asp	gtt Val	ttc Phe	gag Glu	432
agg Arg 145	aca Thr	cgg Arg	cgg Arg	ccg Pro	agg Arg 150	gct Ala	aag Lys	gcg Ala	cca Pro	ctg Leu 155	aga Arg	gtc Val	tcc Ser	gtt Val	tat Tyr 160	480
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caa Gln	att Ile	ttg Leu 195	cga Arg	gcc Ala	ggc Gly	tcg Ser	ggt Gly 200	gtg Val	cgt Arg	ata Ile	tgt Cys	ggc Gly 205	ctt Leu	cct Pro	gat Asp	624
ccc Pro	aaa Lys 210	cgt Arg	ccc Pro	gga Gly	cac His	ctg Leu 215	tgt Cys	tgt Cys	gcc Ala	gat Asp	aat Asn 220	cca Pro	ttg Leu	acg Thr	tgt Cys	672
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acg Thr	gag Glu	tct Ser	gga Gly 260	atc Ile	tgt Cys	gtg Val	aaa Lys	aac Asn 265	ctg Leu	gaa Glu	gaa Glu	cgc Arg	aac Asn 270	Met	acg Thr	816

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										ttg Leu						912
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										tgg Trp						1008
ctg Leu	aaa Lys	ctt Leu	cgc Arg 340	tac Tyr	gtg Val	tgc Cys	aat Asn	gat Asp 345	gat Asp	gtg Val	tca Ser	gat Asp	gat Asp 350	gtc Val	agc Ser	1056
aac Asn	ggt Gly	gcc Ala 355	gcg Ala	gga Gly	gat Asp	gac Asp	agc Ser 360	gly ggg	gac` Asp	gag Glu	gga Gly	ccg Pro 365	tct Ser	gga Gly	gcg Ala	1104
ggt Gly	gtc Val 370	ggt Gly	gct Ala	tcg Ser	gga Gly	aca Thr 375	acg Thr	gga Gly	agc Ser	aca Thr	tct Ser 380	gta Val	tct Ser	acc Thr	ctc Leu	1152
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1				5					10	Asp				15		
			20					25					30			
		35					40			Asp		45				
Asp	Tyr 50	Asn	Lys	Ile	Phe	Asp 55	Asp	Phe	Cys	Ser	Ala 60		Gly	Val	Cys	
Gln 65	Thr	Gly	Ser	His	Ala 70	Gln	Lys	Phe	Lys	Lys 75	Ile	Arg	Met	Leu	Tyr 80	
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Ala	Gly	Gly	Val		Gly	Glu	Arg	Tyr 105		Leu	Phe	Gln	Leu 110		Pro	
Glu	Val			Gly	Cys	Asp			Asn	Leu	Ile	Ala 125	Thr	Thr	Ser	
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	130 Thr	Arg	Arg	Pro		135 Ala	Lys	Ala	Pro		140 Arg		Ser	Val	Tyr	
145 Lys	Arg	Lys	Ser	Lys	150 Arg	Leu	Gln	Asp	Ser	155 Ser	Ala	Gln	Pro	Val	160 Leu	
-	_	-		165	_				170					175		

Gly Ala Val Glu Val Ser Phe Phe Tyr Phe Gly Glu Asn Val Gly Val 185 Gln Ile Leu Arg Ala Gly Ser Gly Val Arg Ile Cys Gly Leu Pro Asp 195 200 205 Pro Lys Arg Pro Gly His Leu Cys Cys Ala Asp Asn Pro Leu Thr Cys 210 215 220 Phe Leu Pro Ser Ser Gln Leu Ile Pro Cys Glu Phe Ala Arg Ala Asp 230 235 Leu Gln Ala Leu Gln Lys Thr Cys Glu Arg Gly Leu Ile Cys Val Met 245 250 255 Thr Glu Ser Gly Ile Cys Val Lys Asn Leu Glu Glu Arg Asn Met Thr 260 265 Ala Leu Thr Asn Tyr Ser Glu Asn Tyr Tyr Glu Leu Arg Pro Ser Gln 275 280 285 Pro Leu Gln Ala Phe Asp Leu Leu His Tyr Leu Arg Glu Leu Ala Arg 295 300 Ser Pro Thr Pro Gly Asp Val Pro Pro Arg Asp Cys Ala Trp Ile Phe 305 310 315 Met Cys Pro Ser Thr Gln Ser Glu Asn Thr Trp Asp Ala Pro Ile Ala 325 330 Leu Lys Leu Arg Tyr Val Cys Asn Asp Asp Val Ser Asp Asp Val Ser 340 345 350 Asn Gly Ala Ala Gly Asp Asp Ser Gly Asp Glu Gly Pro Ser Gly Ala 360 365 355 Gly Val Gly Ala Ser Gly Thr Thr Gly Ser Thr Ser Val Ser Thr Leu 370 380 375 Ala Pro Tyr Gly Arg Lys 385 390

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agt Ser	cag Gln	acc Thr	gtc Val 100	gaa Glu	gat Asp	gta Val	tcc Ser	act Thr 105	gag Glu	gag Glu	aac Asn	ctg Leu	tcg Ser 110	gcg Ala	ccg Pro	336
gcg Ala	cct Pro	aac Asn 115	agg Arg	tgc Cys	cgc Arg	gtt Val	att Ile 120	cgc Arg	ctg Leu	ttg Leu	ccg Pro	atc Ile 125	ttt Phe	gta Val	cga Arg	384
tct Ser	tgc Cys 130	ccg Pro	ctc Leu	tgt Cys	aac Asn	gaa Glu 135	gcg Ala	gat Asp	gcc Ala	acc Thr	ggc Gly 140	ggc Gly	atg Met	ctc Leu	ctg Leu	432
gac Asp 145	gta Val	cgc Arg	aac Asn	gag Glu	gta Val 150	acc Thr	gcc Ala	aga Arg	ttc Phe	cgg Arg 155	tat Tyr	ctc Leu	ggt Gly	gcc Ala	999 Gly 160	480
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										ttg Leu						576
										ccg Pro						624
ctg Leu	caa Gln 210	gga Gly	cac His	gtt Val	tgt Cys	gcg Ala 215	ggc Gly	att Ile	cgg Arg	ccg Pro	gaa Glu 220	cag Gln	gcc Ala	ctg Leu	ttg Leu	672
ccg Pro 225	cat His	acc Thr	cca Pro	cag Gln	gat Asp 230	atg Met	ttt Phe	cct Pro	cac His	cag Gln 235	acg Thr	agc Ser	atg Met	cta Leu	aag Lys 240	720
tgg Trp	ctg Leu	gly 333	aag Lys	gag Glu 245	atc Ile	ata Ile	cgc Arg	999 999	ttg Leu 250	atg Met	att Ile	tac Tyr	gca Ala	gac Asp 255	Gly 999	768
tct Ser	gly ggg	att Ile	tac Tyr 260	att Ile	cgg Arg	tat Tyr	atg Met	ggt Gly 265	cac His	gtt Val	cca Pro	gcc Ala	ttc Phe 270	ctg Leu	ctg Leu	816
ggt Gly	aac Asn	gga Gly 275	ggt Gly	tcg Ser	ctg Leu	gag Glu	ccg Pro 280	gtg Val	gat Asp	ata Ile	att Ile	aac Asn 285	aac Asn	gcg Ala	cga Arg	864
										ctg Leu						912
acc Thr 305	ccg Pro	cca Pro	cac His	gga Gly	acg Thr 310	cga Arg	ttt Phe	cca Pro	gcc Ala	gcc Ala 315	tat Tyr	gcg Ala	tct Ser	ctc Leu	cac His 320	960
cta	gga	ggc	gtt	ccc	act	ccg	gaa	ggc	gag	ccg	tgt	ccc	aca	atc	ccc	1008

Leu Gly Gly Val Pro Thr Pro Glu Gly Glu Pro Cys Pro Thr Ile Pro ctq tcc att caa att tqq cac qaq tgt ctg tgg cgg gcg tgc ggg gat Leu Ser Ile Gln Ile Trp His Glu Cys Leu Trp Arg Ala Cys Gly Asp gcg gcc cag tga Ala Ala Gln <210> 127 <211> 355 <212> PRT <213> Macaca mulatta rhadinovirus 17577 <400> 127 Met Ala Glu Gly Arg Ala Gly Ser Ile Arg Val Asn Arg Pro Ser Gly Leu Arg Ala Trp Leu Leu Asp Cys Cys Asp Asn Asp Lys His Pro Gly Met His Trp Leu Asp Glu Glu Lys Thr Leu Val Arg Leu Pro Trp Asn His Leu Lys Gly Ala Gly Gly Val Ser Asp Asp Glu Arg Asn Met Tyr Leu Asp Tyr Cys Gln Phe Lys Gly Ile Arg Gln Thr Gly Asn Arg Arg Leu Ser Val Arg Glu Cys Lys Asn Trp Leu Ala Ser Ala Ile Arg His Ser Gln Thr Val Glu Asp Val Ser Thr Glu Glu Asn Leu Ser Ala Pro Ala Pro Asn Arg Cys Arg Val Ile Arg Leu Leu Pro Ile Phe Val Arg Ser Cys Pro Leu Cys Asn Glu Ala Asp Ala Thr Gly Gly Met Leu Leu Asp Val Arg Asn Glu Val Thr Ala Arg Phe Arg Tyr Leu Gly Ala Gly Met Glu Tyr Glu Gly Ala Val Gly Gly Asp Gly Glu Gln Cys Trp Met Leu Arg Leu Val Val Tyr Tyr Gly Arg Leu Val Gly Asn Met Glu Val Gly Ser Pro Asn Gly Val Arg Leu Leu Pro Ala Pro Lys Arg Pro Leu Gln Gly His Val Cys Ala Gly Ile Arg Pro Glu Gln Ala Leu Leu Pro His Thr Pro Gln Asp Met Phe Pro His Gln Thr Ser Met Leu Lys Trp Leu Gly Lys Glu Ile Ile Arg Gly Leu Met Ile Tyr Ala Asp Gly Ser Gly Ile Tyr Ile Arg Tyr Met Gly His Val Pro Ala Phe Leu Leu Gly Asn Gly Gly Ser Leu Glu Pro Val Asp Ile Ile Asn Asn Ala Arg Val Leu Arg Val Phe Ser Leu Ala Gln Tyr Leu Ser Ala Val Ser Ala Thr Pro Pro His Gly Thr Arg Phe Pro Ala Ala Tyr Ala Ser Leu His Leu Gly Gly Val Pro Thr Pro Glu Gly Glu Pro Cys Pro Thr Ile Pro Leu Ser Ile Gln Ile Trp His Glu Cys Leu Trp Arg Ala Cys Gly Asp

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350

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345

340

Ala Ala Gln

Ile Thr Pro	Phe Trp 180	Arg Leu	Arg	Ile 185	Arg	Val	Phe	Tyr	Phe 190	Gly	Ser	
ctg gta gcg Leu Val Ala 195												624
aag cgc caa Lys Arg Gln 210	gac ccc Asp Pro	aaa ccg Lys Pro 215	Gly	cac His	gag Glu	tgt Cys	ttc Phe 220	tat Tyr	ggg Gly	aca Thr	gcg Ala	672
tat aag atg Tyr Lys Met 225												720
gag cag aga Glu Gln Arg	gaa acc Glu Thr 245	gtg tgt Val Cys	gag Glu	att Ile	ata Ile 250	aac Asn	999 Gly	tgt Cys	gag Glu	gag Glu 255	ggc Gly	768
gtc ttt ttg Val Phe Leu	cat ggc His Gly 260	aat gag Asn Glu	ctg Leu	ggg Gly 265	atg Met	tat Tyr	gtg Val	gat Asp	aac Asn 270	aga Arg	acc Thr	816
agg cac acg Arg His Thr 275	Val Arg	tgc gca Cys Ala	999 Gly 280	aac Asn	gac Asp	gca Ala	gag Glu	ggg Gly 285	aac Asn	cac His	gca Ala	864
caa cgg gct Gln Arg Ala 290	gtg cga Val Arg	tcc tct Ser Ser 295	Val	aaa Lys	tct Ser	caa Gln	atc Ile 300	ttc Phe	tat Tyr	gtt Val	atg Met	912
ggt cta ctg Gly Leu Leu 305	cgc aga Arg Arg	ctc gcc Leu Ala 310	cgg Arg	tca Ser	ccc Pro	gtt Val 315	ccc Pro	ggc Gly	gac Asp	act Thr	gtt Val 320	960
ccc agc aac Pro Ser Asn												1008
aaa aga ccc Lys Arg Pro	cag gtc Gln Val 340	cct gto Pro Val	act Thr	ttg Leu 345	gtg Val	atc Ile	tgt Cys	cag Gln	gat Asp 350	gaa Glu	ttg Leu	1056
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Ser Asn Leu	20			25					30			
Glu Glu Lys 35		Phe Arg	ı Ile 40	Ser	Trp	His	Arg	Gly 45	Met	Ser	Gly	
Met Gln Pro	Val Val	Ala Tyr	Cys	Leu	Asp	Arg	Asp	Leu	Glu	Cys	Gly	

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Arg Glu Asn Ala Gly Phe Glu Gln Asp Asp Ala Arg Ala Thr Thr Thr
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               85
Arg Phe Gly Glu Arg Phe Phe Tyr Leu Arg Pro Ala Val Asp Pro
                       105
           100
Leu Cys Tyr Ala Cys Ile Leu Asp Ser His Ser Glu Thr Val Leu Asn
                                 125
                120
     115
Tyr Leu Glu Ala Ala Cys Val His Gly Leu Glu Pro Gly Thr Pro Leu
                     135
Pro Pro Pro Ala Pro Ala Glu Ala Asp Gly Ala Ala Arg Ser Val Tyr
                                     1.55
                  150
145
Ala Arg Ala Ala Arg Leu Ala Thr Val Ala Pro Pro His Pro Asp Gln
              165
                                 170
                                                   175
Ile Thr Pro Phe Trp Arg Leu Arg Ile Arg Val Phe Tyr Phe Gly Ser
                                                190
                             185
           180
Leu Val Ala Glu His Thr Ser Gln Asp Arg Arg Gly Val Arg Leu His
                         200
Lys Arg Gln Asp Pro Lys Pro Gly His Glu Cys Phe Tyr Gly Thr Ala
                                        220
                     215
Tyr Lys Met Trp Leu Pro Lys Pro Gln Leu Asp Gly Pro Leu Thr Pro
                                     235
                230
225
Glu Gln Arg Glu Thr Val Cys Glu Ile Ile Asn Gly Cys Glu Glu Gly
                                250
              245
Val Phe Leu His Gly Asn Glu Leu Gly Met Tyr Val Asp Asn Arg Thr
                             265
Arg His Thr Val Arg Cys Ala Gly Asn Asp Ala Glu Gly Asn His Ala
                          280
                                         285
      275
Gln Arg Ala Val Arg Ser Ser Val Lys Ser Gln Ile Phe Tyr Val Met
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Gly Leu Leu Arg Arg Leu Ala Arg Ser Pro Val Pro Gly Asp Thr Val
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Pro Ser Asn Ala Val Thr Leu Tyr Leu Gly Gly Arg Pro Gly Ser Ser
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Lys Arg Pro Gln Val Pro Val Thr Leu Val Ile Cys Gln Asp Glu Leu
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Ser Gly Thr Val Ser Ala Ser Pro Phe Ile Leu Cys Phe Ile Tyr His
            2.0
                                2.5
tog otg tat tit gia gag occ otg att ago git gag aac att ata tic 144
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		cta Leu														240
atc Ile	tcg Ser	gcg Ala	ttt Phe	ctt Leu 85	att Ile	acc Thr	gcc Ala	ggt Gly	tct Ser 90	atg Met	gca Ala	tcc Ser	acc Thr	ctc Leu 95	ggc Gly	288
gtt Val	gac Asp	ctt Leu	cca Pro 100	tgg Trp	gtt Val	cac His	gtt Val	tcc Ser 105	att Ile	ttt Phe	gtg Val	999 999	tcg Ser 110	tgc Cys	ctg Leu	336
		ctg Leu 115														384
ccc Pro	acg Thr 130	att Ile	gcc Ala	cac His	aga Arg	tac Tyr 135	tac Tyr	gaa Glu	ctt Leu	gga Gly	ttt Phe 140	tta Leu	gca Ala	gcg Ala	ttc Phe	432
		tat Tyr														480
ttt Phe	ttg Leu	ctg Leu	ccc Pro	ctg Leu 165	gtg Val	gcc Ala	ttt Phe	ata Ile	gta Val 170	ggt Gly	ggc Gly	gtt Val	tgt Cys	tca Ser 175	ctc Leu	528
		ctg Leu														576
		att Ile 195														624
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ttg Leu 225	att Ile	gga Gly	999 Gly	gcg Ala	gcc Ala 230	gcg Ala	ggt Gly	acg Thr	ctg Leu	tct Ser 235	gtt Val	ggc Gly	ctg Leu	acg Thr	acg Thr 240	720
		ctc Leu														768
		tgt Cys														816
gtt Val	tat Tyr	gtt Val	tta Leu	gca Ala	gcc Ala	gct Ala	gtg Val	ctg Leu	ctg Leu	acg Thr	ctc Leu	acg Thr	cac His	gtc Val	ttg Leu	864

280 285 275 ggg cca gga acg cat aat ttg ttc acc aga gtg tgt gtg ttt acg gtt Gly Pro Gly Thr His Asn Leu Phe Thr Arg Val Cys Val Phe Thr Val 290 295 300 ttt tta ttg act atg ttt ggg gcg att gga tgc gaa tta caa ata atc Phe Leu Leu Thr Met Phe Gly Ala Ile Gly Cys Glu Leu Gln Ile Ile 310 aga aaa aaa cta cag cgt gcc gcg aac tcg cca aga ata gtc ttg ggg 1008 Arg Lys Lys Leu Gln Arg Ala Ala Asn Ser Pro Arg Ile Val Leu Gly gtg tgt gcc tgc gga aac ctt ctg atg gcg gtg gtt ttt ttc tcc tta Val Cys Ala Cys Gly Asn Leu Leu Met Ala Val Val Phe Phe Ser Leu 340 345 1083 aat aaa gtt gag ctt ggt gcc ctt taa Asn Lys Val Glu Leu Gly Ala Leu 355 <210> 131 <211> 360 <212> PRT <213> Macaca mulatta rhadinovirus 17577 <400> 131 Met Gly Thr Tyr Thr Ser Glu Ala Ser Leu Ala Trp Leu Ser Phe Met 10 5 Ser Gly Thr Val Ser Ala Ser Pro Phe Ile Leu Cys Phe Ile Tyr His 20 25 Ser Leu Tyr Phe Val Glu Pro Leu Ile Ser Val Glu Asn Ile Ile Phe 45 40 Ser Trp Gly Ala Val Gly Leu His Gly Leu Leu Leu Leu Phe Cys Ile 55 Phe Gly Leu Pro Ala Trp Leu Ser Arg Gln Val Asp Val Pro Cys Thr 70 75 Ile Ser Ala Phe Leu Ile Thr Ala Gly Ser Met Ala Ser Thr Leu Gly 85 90 Val Asp Leu Pro Trp Val His Val Ser Ile Phe Val Gly Ser Cys Leu 105 110 Cys Leu Leu Leu Cys Val Val Ala Ala Asn Asp Val Val Tyr Leu Cys 125 120 Pro Thr Ile Ala His Arg Tyr Tyr Glu Leu Gly Phe Leu Ala Ala Phe 140 135 Ser Val Tyr Tyr Phe Leu Val Leu Lys Asn Leu Phe Leu Ala Pro Val 150 155 Phe Leu Leu Pro Leu Val Ala Phe Ile Val Gly Gly Val Cys Ser Leu 170 165 Arg Ala Leu Arg Ser His Pro Leu Tyr Glu Ala Gly Leu Gln Arg Arg 190 180 185 His Ala Ile Phe Ser Leu Thr Ser Arg Arg Tyr Ile Thr Tyr Ser Ile 200 205 Lys Gln Ala Leu Glu Val Cys Gly Trp Asp Phe Tyr Leu Val Thr Val 220 215 Leu Ile Gly Gly Ala Ala Ala Gly Thr Leu Ser Val Gly Leu Thr Thr 230 235 Pro Leu Leu Leu Gly Leu Val His Tyr Phe Phe Val Phe His Val Gly. 255 250

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gca ctg gcg ccg ctg atg gta tat tct gac atg acg gac gag gtt agc 240
Ala Leu Ala Pro Leu Met Val Tyr Ser Asp Met Thr Asp Glu Val Ser
65 70 75 80

ttt agc ttt cga aac acc tcc ctt ggg aac acg ttc aca cac acc cgt 288
Phe Ser Phe Arg Asn Thr Ser Leu Gly Asn Thr Phe Thr His Thr Arg
85 90 95

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gag Glu	tct Ser 770	aac Asn	tgc Cys	gta Val	caa Gln	aag Lys 775	gcc Ala	gac Asp	ggt Gly	gag Glu	cgg Arg 780	acc Thr	aag Lys	gta Val	tgt Cys	2352
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4.0 Gly Trp Asp Ile Glu Ala Asn Ser Leu Thr Gly Leu Leu Trp His Arg Ile Met Glu Asp Arg Cys Leu Val Thr Val Arg Asp Tyr Leu Ala Val 7.0 Phe Gly Glu Arg Leu Ser Asp Glu Val Arg Ala Phe Met Ser Lys His Glu Ala Ala Leu Asp Gly Leu Leu Gln Asp Phe Lys Gln Ser Lys Ala Tyr Thr Asn Phe Val Asn Cys Gly Tyr Leu Ser Ala Val Arg Phe Tyr Asp Thr Tyr Val Leu Arg Thr Gln Gly Ser Ser Pro Ile Phe Glu Ser Val Ala Gln Met Phe Met Arg Val Ala Val Phe Val Ala Cys Gln Cys Ile Lys Phe Pro Cys Leu Arg Glu Thr Leu Arg His Leu Val Glu Ser Glu Thr Glu Leu Asp Glu Met Tyr Leu Val Gly Tyr Ala Phe His Tyr Ile Ser Ser Gln Ile Val Cys Cys Ala Thr Pro Val Leu Arg Ser Ala Gly Leu Arg Gly Gly Gln Leu Ser Ser Cys Phe Ile Leu Lys Pro Ser Met Ala Thr Glu Asp Lys Thr Leu Lys Ala Leu His Glu Glu Met Ser Pro Leu Leu Ala Ser Lys Ser Gly Val Gly Ile Asp Val Ser Ser Phe Ala Glu His Lys Asn Ile Thr Ser Cys Leu Lys Leu Ile Asn Ala His Val Gly Tyr Phe Asn Asp Asn Asn Ile Arg Pro Val Gly Ala Ser Ala Tyr Met Glu Leu Trp His His Gln Ile Cys Asp Phe Leu Asn Ala Lys Met Pro Glu Asn Gln Glu Arg Cys His Asn Leu Phe Gln Gly Val Cys Val Pro Glu Leu Phe Phe Arg Leu Tyr Glu Thr Asn Pro Asp Gly Gln Trp His Leu Phe Ala Pro Glu Val Ala Pro Asn Leu Leu Lys Leu Tyr Gly Ala Glu Phe Glu Ile Glu Tyr Asn Arg Leu Val Ala Ala Gly Lys His Ser Ser Ser Leu Pro Leu Lys Ser Met Met Tyr Ala Leu Ile Asn Thr Val Ile Lys Thr Gly Ser Pro Tyr Val Leu Leu Lys Glu Ala Leu Asn Lys His His Trp Cys Glu Thr Gln Gly Ser Ala Ile Asn Cys Ser Asn Leu Cys Ala Glu Ile Val Gln Gln Pro Glu Gly Gln Ala Ser Val Cys Asn Leu Ala Asn Ile Ser Leu Pro Lys Cys Leu Arg Pro His Arg Gly Glu Ser Gly Val Glu Pro Gly Lys Gly Asp Val Thr Phe Gly Phe Glu Leu Leu Asp Asp Ala Val Glu Ala Ala Val Ile Ile Val Asn Ala Cys Ile Leu Gly Gly Thr Ala Pro Thr Glu Ser Val Arg Arg Gly Gln Glu Glu Arg Ser Met Gly Ile Gly Val Gln Gly Leu Ala Asp Val Phe Ala Glu Leu Gly Phe Gly Tyr Leu Asp Ala Glu Ser Ala Lys Leu Asp 

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	acg Thr												912
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	aag Lys												1200
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	acg Thr															1968
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	ccc Pro				_			_								2496
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	gcc Ala															2688
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395

415

Thr Pro Lys Trp Glu Asn Phe Thr Asp Phe Leu Glu Met Val Asn Gln

Leu Thr Leu Tyr Gly Phe Tyr Phe Tyr Glu Cys Leu Asn Gln Tyr Ser

390

Pro Thr Ser Ile Ser Leu Ala Lys Ile Gln Asn Ile Leu Asn Arg Val Asp Ala Glu Gln Ser Asp Arg Ala Leu Trp Arg Thr Pro Leu Ile Gly Ser Phe Pro Phe Pro Trp Lys Leu Asn Asn Val Leu Ala Phe Phe Lys Pro Ser Thr Pro Val Ala Thr Leu Gln Lys Ile Tyr Lys Ala Ile Pro Ser Tyr Leu Met Arg Ser Leu Phe Glu Ile Ala Ala Asn Lys Ser Trp Gly Asn Ile Ala Leu Ala Glu Ser Ala Pro Leu Thr Asp Ile Gln Thr Ala Glu Pro Asp Gln Gly Pro Val Ser Ala Gln Val Ile Ala Lys Tyr Cys Ser Arg Leu Gln Ile Ser Ala Thr Asp Tyr Asp Ala Ala Ile Val Ser Ser Pro Gly Phe Ala Ala Glu Phe Ile Lys Thr Lys Leu Tyr Pro Ile Leu Ser Glu Val Leu Arg Asn Thr Ser Lys Lys Asn Arg Ser Leu Phe Gln Ile Arg Trp Leu Ile Val Phe Ala Ala Glu Asp Ala Arg Asp Leu Ala Pro Ile Arg Arg Ser Leu Ala Leu Ala Tyr Phe Gln Ile Met Asp Ile Leu Glu Glu Lys His Ser Pro Glu Ser Phe Tyr Asn Leu Leu Asp Tyr Leu Gln Glu Thr Phe Arg Cys Ile Arg Gln Val Ile Pro Glu Ala Thr Cys Pro Gln Glu Phe Leu Gln Tyr Leu Phe Thr Phe Gln Asn Ile Pro Ile Ala Ala Ser Phe Ile Gln Thr Ser Met Thr Phe Val Asp Asp Leu Lys Asn Gly Ile Pro Gly Ile Leu Asp Leu Val Ser Leu Gly Ala Ala Phe Tyr Asn Met Lys Leu Leu Tyr Asp Ser Thr Leu Asp Thr Val Glu Ile Pro Thr Glu Glu Gly Gln Pro Ile Val Val Ser Met Phe Val Phe Lys Ser Thr Ile Arg Val Leu Glu Lys Leu Leu Gln Glu Ala Val Ile Ala Leu Thr Gln Thr Ser Glu Pro Met Tyr Ala Ala His Ile Arg Leu Met Gln His Leu Thr Tyr Met Gln Lys Ile Ala Gly His Glu Ile Met Thr Thr Gln Leu Pro Ser Val Phe His Glu Ile His Glu Gly Tyr Leu Gln Cys Phe Lys Arg Phe Lys Arg Leu Met Leu His Val Thr Gly Ser Cys Cys Tyr Ser Leu Thr Arg Tyr Phe Gly Phe Leu Tyr Gln Pro Pro Leu Ile Pro Asp Thr Ile Val Gln Lys Ile Leu Asn Phe Asn Asp Lys Thr Asp Thr Thr Asp Asp Ile Leu Lys Ser Leu Ser Gln Pro Val Arg Gln Gly Pro Leu Ser Ala Glu Asn Glu Ser Ser Ser Arg Leu Ser Lys Asn Asn Val Glu Leu Leu Gln Lys Leu Tyr Asp Asp Phe Arg Thr Ala Ser Thr Asn Asn Asn Pro Thr Ser Ile Lys Leu Glu Tyr Ser Gly Asn Tyr Asn Glu Thr Gln Val Ser Val Asp Trp Ser Thr Tyr Asn Leu Val Thr Tyr Thr Ala Pro Asp Asp Thr Leu Lys Phe Thr Pro Val
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		_					_		-	_		-	atg Met 750			2256
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													gaa Glu			2352
cca	cca	att	C 2 2	~~~	~-~											2400
	_	_	_		_						_		gca Ala			2400
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Pro 785 gaa Glu ctg	Pro tcg Ser	Val gta Val gac	Gln aca Thr	Glu aaa Lys 805 gaa	Leu 790 atg Met	Pro gaa Glu tta	Leu aag Lys ctt	Val aac Asn	Ala caa Gln 810 aca	Arg 795 cag Gln ata	Ala gct Ala acc	Lys ctc Leu gca	Ala gac	caa Gln 815	Met 800 ata Ile gga	
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Pro 785 gaa Glu ctg Leu gat Asp	tcg ser gga Gly gag Glu	yal gta val gac Asp aac Asn 835	aca Thr gcc Ala 820 ccg Pro	aaa Lys 805 gaa Glu gtc Val	Leu 790 atg Met acg Thr cgc Arg	gaa Glu tta Leu gcc Ala	aag Lys ctt Leu atg Met 840	val aac Asn gac Asp 825 tcc ser	caa Gln 810 aca Thr ata Ile	Arg 795 cag Gln ata Ile ccg Pro	gct Ala acc Thr ata Ile	ctc Leu gca Ala ctg Leu 845	gac Asp aca Thr 830	caa Gln 815 tcc Ser acc Thr	Met 800 ata Ile gga Gly tac Tyr	2448 2496
Pro 785 gaa Glu ctg Leu gat Asp att Ile	tcg ser gga Gly gag Glu aca Thr 850	yal gta val gac Asp aac Asn 835 aac	aca Thr gcc Ala 820 ccg Pro gca Ala	aaa Lys 805 gaa Glu gtc val	Leu 790 atg Met acg Thr cgc Arg	gaa Glu tta Leu gcc Ala ctg Leu 855	aag Lys ctt Leu atg Met 840 ata Ile	val aac Asn gac Asp 825 tcc ser ggc Gly	caa Gln 810 aca Thr ata Ile agt Ser	Arg 795 cag Gln ata Ile ccg Pro tct ser	Ala gct Ala acc Thr ata Ile cga Arg 860 tca	ctc Leu gca Ala ctg Leu 845 aac	gac Asp aca Thr 830 gag Glu	caa Gln 815 tcc Ser acc Thr	Met 800 ata Ile gga Gly tac Tyr ttc Phe	2448 2496 2544
Pro 785 gaa Glu ctg Leu gat Asp att Ile gaa Glu 865	tcg ser gga Gly gag Glu aca Thr 850 aaa Lys	yal gta val gac Asp aac Asn 835 aac Asn ctc Leu atg	Gln aca Thr gcc Ala 820 ccg Pro gca Ala aag Lys	aaa Lys 805 gaa Glu gtc Val ggc Gly	Leu 790 atg Met acg Thr cgc Arg gcc Ala gcc Ala 870 aac	gaa Glu tta Leu gcc Ala ctg Leu 855 atc	aag Lys ctt Leu atg Met 840 ata Ile cac	aac Asn gac Asp 825 tcc ser ggc Gly gac Asp	caa Gln 810 aca Thr ata Ile agt Ser ctg Leu	Arg 795 cag Gln ata Ile ccg Pro tct ser gca Ala 875 gat	gct Ala acc Thr ata Ile cga Arg 860 tca Ser	ctc Leu gca Ala ctg Leu 845 aac Asn	gac Asp aca Thr 830 gag Glu cag Gln	caa Gln 815 tcc Ser acc Thr cgg Arg	Met 800 ata 11e gga Gly tac Tyr ttc Phe 880 aat	2448 2496 2544 2592

	900	905		910
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cgc gaa tgc Arg Glu Cys 930	gta gaa gcg Val Glu Ala	cta aat aaa Leu Asn Lys 935	agg agc ccc tct Arg Ser Pro Ser 940	tcc ctc aac 2832 Ser Leu Asn
aac gcg cgt Asn Ala Arg 945	ctc ctc gcg Leu Leu Ala 950	gtt caa acc Val Gln Thr	ata ctg ggg cac Ile Leu Gly His 955	gcg tcc gtt 2880 Ala Ser Val 960
cca gat cac Pro Asp His	gag acg ctg Glu Thr Leu 965	acg cga atc Thr Arg Ile	gtt tcc ggc gtc Val Ser Gly Val 970	gcc agc gca 2928 Ala Ser Ala 975
Gln Lys Glu	tcc gct ggc Ser Ala Gly 980	gat gat cca Asp Asp Pro 985	gat agg tgg acg Asp Arg Trp Thr	cga gta acc 2976 Arg Val Thr 990
ggt cac cta Gly His Leu 995	aac gag ctg Asn Glu Leu	aag ctc gta Lys Leu Val 1000	act acc caa tcg Thr Thr Gln Ser 1005	cgt gtc gac 3024 Arg Val Asp
aaa gcc acc Lys Ala Thr 1010	Arg Arg Lys	ctg tta atg Leu Leu Met 015	ata ata acc cgt Ile Ile Thr Arg 1020	gac ctc aag 3072 Asp Leu Lys
gag gcg gag Glu Ala Glu 1025	gtg tct cag Val Ser Gln 1030	gaa acg gtc Glu Thr Val	ctg gaa aca cgg Leu Glu Thr Arg 1035	tgg caa gaa 3120 Trp Gln Glu 1040
aac gtg cta Asn Val Leu	aag ttt caa Lys Phe Gln 1045	Pro Ser Thr	tcc aaa gaa atc Ser Lys Glu Ile 1050	gaa gac ttt 3168 Glu Asp Phe 1055
Leu Gln Ser	gca ccg tca Ala Pro Ser 060	gca aag gcc Ala Lys Ala 1065	cga aaa ttc gca Arg Lys Phe Ala	gaa aaa cac 3216 Glu Lys His 1070
cta cgg acg Leu Arg Thr 1075	ctg atc acc Leu Ile Thr	caa ttc aac Gln Phe Asn 1080	ggc cac gag cga Gly His Glu Arg 1085	ccg ccg tcc 3264 Pro Pro Ser
gag gcc acc Glu Ala Thr 1090	Ala Val Pro	atg gac tac Met Asp Tyr 095	acg ccg acg ccc Thr Pro Thr Pro 1100	ata ccc acg 3312 Ile Pro Thr
cca cag gcc Pro Gln Ala 1105	gtt tct acg Val Ser Thr 1110	gct acc gcg Ala Thr Ala	gaa aag gga aag Glu Lys Gly Lys 1115	gcc gca tgg 3360 Ala Ala Trp 1120
aat aaa att Asn Lys Ile	caa cag gcc Gln Gln Ala 1125	Phe Gln Asp	ttc aac ttt cac Phe Asn Phe His 1130	ctc atc gac 3408 Leu Ile Asp 1135
Ala Ser Asp	tgg caa gag Trp Gln Glu 140	atg gca tca Met Ala Ser 1145	gaa tac tcc aga Glu Tyr Ser Arg	cac ggc tcg 3456 His Gly Ser 1150

tcc ctt cct ggt Ser Leu Pro Gly 1155	Thr Val Gly P	cca aag ctg Pro Lys Leu 160	gtg cgc ttc at Val Arg Phe Me 1165	g gag agc t Glu Ser	3504
atc tca aac acc Ile Ser Asn Thr 1170	ctg gac gac a Leu Asp Asp I 1175	atc ctc acg [le Leu Thr	cag aag ctg go Gln Lys Leu Al 1180	a tct ctg a Ser Leu	3552
ctt cca aac ggg Leu Pro Asn Gly 1185	ccc gcg ttc a Pro Ala Phe A 1190	Arg Pro Pro	gcg ttt gac to Ala Phe Asp Tr 195	gg atc gcg pp Ile Ala 1200	3600
ccc tat caa aca Pro Tyr Gln Thr	cgc gta aac o Arg Val Asn A 1205	gcg ttt cta Ala Phe Leu 1210	aaa acc ata gg Lys Thr Ile Gl	gc ctg ccc Ly Leu Pro 1215	3648
atg gtg cgc aac Met Val Arg Asn 1220	ctg gcg gac a Leu Ala Asp I	aag atc cat Lys Ile His 1225	cac caa tgc ca His Gln Cys Gl 123	ln Thr Val	3696
agt cac gcg gtg Ser His Ala Val 1235	Gln Ser Ala A	gac ctt caa Asp Leu Gln 240	cag gcc acg gt Gln Ala Thr Va 1245	g gga aca al Gly Thr	3744
agt tta gaa cga Ser Leu Glu Arg 1250	ccc gcg gcc g Pro Ala Ala ( 1255	gaa tac tgt Glu Tyr Cys	cga ata ctc to Arg Ile Leu So 1260	ct gac atg er Asp Met	3792
caa gtc gcg ttc Gln Val Ala Phe 1265	aac gac cac g Asn Asp His 0 1270	Gly Ile Ala	gta aga tcg ga Val Arg Ser G 275	ag gcc gcg lu Ala Ala 1280	3840
gcg tac acg gac Ala Tyr Thr Asp	gca atc aac t Ala Ile Asn S 1285	tcg ccg gcc Ser Pro Ala 1290	aac gtc gtg a Asn Val Val T	ct ccc ccg hr Pro Pro 1295	3888
aaa ccc aac cta Lys Pro Asn Leu 1300	Glu Ala Pro I	aag aag cta Lys Lys Leu 1305	ata acg gca a Ile Thr Ala T 13	hr Asp Ala	3936
cta acc gtc gag Leu Thr Val Glu 1315	Asp Phe Pro	gat ttc cta Asp Phe Leu 320	aaa acg tca a Lys Thr Ser I 1325	tc ctt caa le Leu Gln	3984
cag gag cag cga Gln Glu Gln Arg 1330	ctc att gcg ( Leu Ile Ala 1 1335	ctc cag aga Leu Gln Arg	gcg gaa ttt c Ala Glu Phe G 1340	ag caa cta ln Gln Leu	4032
gag gcc agc atc Glu Ala Ser Ile 1345	tcg gcg gcc g Ser Ala Ala ( 1350	Glu Arg Leu	cgc caa tcc a Arg Gln Ser T 1355	cc cgt gac hr Arg Asp 1360	4080
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gcc ccc gtc gca Ala Pro Val Ala 1380	Ile Ser Ser	aga ccg ttg Arg Pro Leu 1385	Asn Leu Ser L	aa cct ata ys Pro Ile 90	4176

gac ttt ttg ag Asp Phe Leu Se 1395	r Ser Thr Val	tac gac aaa a Tyr Asp Lys I 400	tc ctg gac aag g le Leu Asp Lys ( 1405	gag cct 4224 Glu Pro
tac gag aca gc Tyr Glu Thr Al 1410	c ata gcg gga a Ile Ala Gly 1415	ttc gcg tgg c Phe Ala Trp L	tg gaa atc gcg a eu Glu Ile Ala ' 1420	aca aaa 4272 Thr Lys
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gat ttg gaa ct Asp Leu Glu Le 146	u Ser Ala Lys	aac acg gac g Asn Thr Asp A 1465	gac gta aag gtg Asp Val Lys Val 1470	ctg aag 4416 Leu Lys
cag gcg cta ga Gln Ala Leu As 1475	p Glu Leu Ala	ccc ctc agg c Pro Leu Arg V 480	gta aag ggc gga Val Lys Gly Gly 1485	aaa acc 4464 Lys Thr
acc gta gac gc Thr Val Asp Al 1490	g tgg aaa caa a Trp Lys Gln 1495	aaa ctg gaa a Lys Leu Glu S	agc ata gaa tcc Ser Ile Glu Ser 1500	ctg ctt 4512 Leu Leu
cgc gcc acg ag Arg Ala Thr Ar 1505	g acg gca ggc g Thr Ala Gly 1510	Glu Ile Ser S	ccg gag ctt gaa Ger Glu Leu Glu 515	cgc atc 4560 Arg Ile 1520
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tcc gat caa tg Ser Asp Gln Cy 154	s Arg Glu Ala	gca aat ttc o Ala Asn Phe I 1545	ctc aga cag gcc Leu Arg Gln Ala 1550	agt cta 4656 Ser Leu
ccc gaa ggc tt Pro Glu Gly Ph 1555	e Ser Asp Ile	ggc aca aaa o Gly Thr Lys 1 1560	ctc agc gag ctt Leu Ser Glu Leu 1565	cag gcg 4704 Gln Ala
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cct aac gtc tt Pro Asn Val Ph 1585	t caa cgc ttc ne Gln Arg Phe 1590	Pro Leu Ser (	caa aac ata acc Gln Asn Ile Thr 595	gaa aac 4800 Glu Asn 1600
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ctt cac gtg cg Leu His Val Ar 162	g Gly Ser Ala	ccc cac ttt a Pro His Phe 1 1625	aca acg tgg ata Thr Thr Trp Ile 1630	gaa acg 4896 Glu Thr
cta ccg acc gt	c gat ccg gaa	aaa cca act	cac gtc ccg gcg	cac gga 4944

Leu I		hr 35	Val	Asp	Pro		Lys .640	Pro	Thr	His		Pro 645	Ala	His	Gly	
gga g Gly 1	-		_		Arg	_		_		Ser		_				4992
ttg t Leu I 1665				Cys			_		Thr	_	_	-	_	Ala		5040
ggt d Gly I			Ile	-			_	Arg	_		_	_	Ala	-		5088
tgg a		sp		_			Asp		_	-		Leu				5136
ctc c	Åsp T	_		_		Thr		_		_	Asn		_		_	5184
ttt a Phe A				_	Ala	_	_	_		Thr						5232
gct a Ala s 1745	_	_	_	Val	_		_	_	Leu	-	-		_	Val		5280
ttt o	_		Glu	_	_			Pro			-		Thr	-	_	5328
acc g Thr A	-	let '				_	Ala	_	-,			Arg				5376
tac t Tyr S	Ser G					Leu	_	-	_		Leu	_		_		5424
cag g Gln A 18		_			Lẹu	_				Gln	_					5472
cat a His A 1825			-	Pro	_	_	-	_	Phe	_				Ala		5520
ccg t Pro T			Pro .				_	Ser					Gln	_	_	5568
tgg g	-	ln T	_	_		_	Gln		_	_		Asn		_	-	5616
gcc a		_			_					_	_		-			5664

1875	1880	1885	
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		acc cta cac ctg acc Thr Leu His Leu Thr 1915	
	o Thr Pro Arg Arg	gaa acg acc acc gag Glu Thr Thr Thr Glu 1930	
		tac tgc atc tcg ggt Tyr Cys Ile Ser Gly 1950	
2 2		ccg gta tcc gct ttc Pro Val Ser Ala Phe 1965	
		ccg atc aga ata ttt Pro Ile Arg Ile Phe 1980	
		cgc ggc ggc atg ggg Arg Gly Gly Met Gly 1995	
	u Cys Val Pro Asp	gtc gag ccc ttc aaa Val Glu Pro Phe Lys 2010	
_		att gaa acg cta ccc Ile Glu Thr Leu Pro 2030	
		ttt ctg aga cag gca Phe Leu Arg Gln Ala 2045	
		gcc gcc cgg tcg tcg Ala Ala Arg Ser Ser 2060	
		aac ctc gta acg gca Asn Leu Val Thr Ala 2075	
	o Ala Asn Phe Glu	agc agg ccg ttt tac Ser Arg Pro Phe Tyr 2090	
		aaa acg ctg tcg gta Lys Thr Leu Ser Val 2110	Thr Ser

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	Thr	acg Thr 2130	cta Leu	tcc Ser	cga Arg	Glu	cac His 2135	gly ggg	acc Thr	gtg Val	Gln	ggc Gly 2140	agg Arg	gat Asp	atc Ile	ttc Phe	6432
	gca Ala 2145	Ala	gct Ala	ccg Pro	Thr	aac Asn 2150	gtc Val	aca Thr	ccg Pro	Glu	caa Gln 2155	acc Thr	gcc Ala	aat Asn	Pro	ccg Pro	6480
•	gca Ala	tgg Trp	gaa Glu	acg Thr	gat Asp 2165	aac Asn	cga Arg	tta Leu	Ile	acg Thr 2170	caa Gln	aca Thr	gaa Glu	Thr	gcc Ala 2175	aaa Lys	6528
	aaa Lys	cct Pro	His	ata Ile 2180	att Ile	cct Pro	gcg Ala	ser	cct Pro 2185	aaa Lys	gcg Ala	cgg Arg	Thr	gat Asp 2190	cca Pro	ccg Pro	6576
	gtg Val	Glu	acc Thr 2195	acg Thr	acc Thr	cac His	His	tca Ser 200	caa Gln	ggg Gly	caa Gln	Ala	tcg Ser 205	caa Gln	cac His	gca Ala	6624
	Asn	agc Ser 2210	Asn	gta Val	aac Asn	Gln	ccc Pro 2215	ggt Gly	caa Gln	att Ile	Thr	tca Ser 2220	cac His	gcg Ala	tca Ser	cgt Arg	6672
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	aac Asn	acg Thr	caa Gln	acg Thr	gtg Val 2245	cct Pro	cga Arg	cta Leu	Ile	tct Ser 2250	caa Gln	acg Thr	tcg Ser	Glu	acg Thr 2255	gcc Ala	6768
	cat His	ata Ile	Asn	cag Gln 2260	cca Pro	gcc Ala	tcc Ser	Gly	cag Gln 2265	gtc Val	acc Thr	gaa Glu	Pro	aag Lys 2270	gga Gly	atc Ile	6816
	ttt Phe	Gly	acg Thr 2275	tat Tyr	aaa Lys	ccc Pro	Arg	gtg Val 280	ctc Leu	acc Thr	gaa Glu	Pro	gcc Ala 2285	aaa Lys	ccc Pro	gca Ala	6864
	Asn	gcc Ala 2290	ggc Gly	gta Val	gcc Ala	Ser	cgc Arg 295	caa Gln	cca Pro	gag Glu	Ala	acc Thr 2300	acc Thr	acg Thr	gtc Val	ccc Pro	6912
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tca ctc acc gtc ccc act ccc aga gtc acc cca atc cct ccc act aac Ser Leu Thr Val Pro Thr Pro Arg Val Thr Pro Ile Pro Pro Thr Asn 2485 2490 2495	7488
atc tgg ata ccc cta tcc cac gtc aac atc caa cac gaa gaa atc aca Ile Trp Ile Pro Leu Ser His Val Asn Ile Gln His Glu Glu Ile Thr 2500 2505 2510	7536
cga gcc aag aat gtg tta atg cga ttt att caa aac gta cga aga aaa Arg Ala Lys Asn Val Leu Met Arg Phe Ile Gln Asn Val Arg Arg Lys 2515 2520 2525	7584
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Cys Leu Ser Asn Cys Val Ile Tyr Leu Ala Gln Ser Tyr Phe Asn Arg 35 40 45	

Glu Ser Pro Val Thr Asp Thr Asn Asp Leu Asp Asp Val Leu Arg Gln Gly Ala Thr Leu Asp Phe Ile Leu Arg Arg Ser Gly Thr Leu Gly Tyr 7.5 Asn Gln Tyr Ala Gln Leu His His Ile Pro Ser Phe Ile Lys Thr Asn Glu Trp Thr Ala Ala Ile Phe Gln Ser Gln Glu Tyr Phe Gly Leu Ile Gly Leu Asp Ala Ala Ile Arg Glu Pro Phe Ile Glu Ser Leu Lys Ser Ile Leu Thr Arg Asn Tyr Ala Gly Thr Val Gln Tyr Phe Leu Phe Ile Cys Gly Asp Lys Ala Gly Ala Val Ile Ile Lys Asn Lys Thr Phe Tyr Leu Phe Asp Pro His Cys Val Pro His Val Pro Asn Ser Pro Ala His Val Ile Ser Ser Ser Asp Pro Thr Ala Ile Leu Glu Tyr Val Ser Pro Pro Asp Arg Glu Tyr Thr Gly Ser Phe Leu Tyr Ile Met Pro Ser Glu Tyr Val Asn Pro Glu His Tyr Ile Thr Asn His Tyr Arg Thr Ile Thr Phe Ala Lys Val His Gly Pro His Ile Asp Ile Ser Thr Gly Ile Glu Pro Cys Thr Ile Glu Asp Ile Pro Ser Pro Pro Arg Ser Pro Asp Val Thr Ser Lys Ser Ser Asn Leu Ala Arg Val Pro Arg Thr Thr Asp Thr Ser Ser Ala Lys Pro Pro Pro Ala Thr Leu Ser Gly Leu Arg Gly Ala Glu Pro Pro Thr Ser Tyr Pro Asp Pro Ala Thr Asn Asp Ala Asp Thr Lys Leu Leu Thr Pro Ala Pro Ala Gln Thr Ala Val Asp His Pro Glu Phe Gln Thr Thr Pro Gly Ala Thr Leu Leu Leu Ser Glu Leu Ser Ala Ser Arg Gly Arg Lys Arg Lys Leu Ser Ser Leu Gln Arg Tyr Ser Asp Ser Asp Glu Ala Ser Ser Asp Asp Glu Gly Ala Pro Arg Arg Arg Val His Asp Asp Ala Ile Ser Ala Glu Val Ile Trp Met Asp Asp Asp Ile Ser Pro Leu Tyr Ser Pro Ser Ala Thr Pro Ser Phe Asp Asp Val Phe Asp Ser Pro Pro Met Ser Pro Glu Phe Thr Tyr Glu Asp Ala Thr Glu Asp Thr Asp Gly Ala Phe Leu Glu Gln Ile Ala Arg Asp Ala Glu Thr Pro Phe Ser Ala Phe Asp Asp Leu Ile Thr Asp His Asp Phe Ser Ser Leu Asp Lys Lys Ile Glu Gln Leu Ile Lys Tyr Glu Ala Pro Ser Gln His Leu Pro Asn Ile Ser Asp Lys Gln Asn Gly Arg Ala Val Arg Glu Ala Ala Ala Leu Gln Ala Met Asp Lys Ile Met Ile Asn Ile Ile Leu Glu His Gly Leu Ile Thr Asp Ala Gln Ala Arg Gly Pro Ser Ala Cys Lys Asn Val Leu Gln Phe Phe Ile Leu Trp Gly Glu Lys Leu Asn Ile Pro Ile Ser Asp Ala Lys Gln Val Leu Glu Leu Asp Leu Gln Leu

	530					535					540				
Ile 545	Pro	Leu	His	Thr	Ala 550	Ile	Ser	Glu	Gly	Lys 555	Phe	Lys	Gln	Gly	Ala 560
Phe	Lys	Lys	His	Leu 565	Thr	Thr	Lys	Ile	Asn 570	Arg	Cys	Leu	Ala	Ser 575	Met
Arg	Ala	Thr	His 580	Ala	Asp	Ala	Gln	Lys 585	Lys	Leu	Ala	Ser	Ala 590	Phe	Asn
Val	Glu	Gly 595	Ser	Gln	Ile	Ser	Ser 600	Ser	Glu	Ala	Lys	Ile 605	Ser	Val	Arg
Ala	Leu 610	Lys	Glu	Gln	Ile	Ala 615	Asn	His	Leu	Ser	Pro 620	Gly	Phe	Leu	Ala
Val 625	Tyr	Ser	Ala	Asp	Glu 630	Val	Lys	His	Leu	Arg 635	Asp	Lys	Ile	Gln	Asp 640
Leu	Lys	Thr	Gly	Ile 645	Glu	Gln	Arg	Asn	Lys 650	Glu	Ile	Gln	Gln	Glu 655	Glu
Leu	Phe	Phe	Asp 660	Ala	Met	Leu	Thr	Ala 665	Leu	Asp	Thr	Phe	Gln 670	Pro	Pro
Pro	Lys	Thr 675	Ala	Phe	Pro	Met	Glu 680	Ile	Phe	Pro	His	Arg 685	Lys	Thr	Glu
	690		_			695					700			Glu	_
705					710		-			715				Gln	720
				725					730					Asn 735	
			740	_				745				_	750	Gln	
		755					760					765		Leu	
-	770					775				_	780	-		Thr	
785					790					795		_		Lys	800
				805			-		810				_	Gln 815	
	-	~	820				-	825					830	Ser Thr	_
_		835					840					845		Arg	
	850					855		-			860			Ser	
865					870					875				Asp	880
				885				J	890	-				895 Leu	
		-	900					905					910	Leu	
		915			-		920				•	925		Leu	
_	930	_				935		_			940			Ser	
945					950					955				Ser	960
	-			965			-		970		-			975 Val	
	•		980		_	-	-	985	-	_	_		990	Val	
•		995					1000					1005		Leu	
-	1010	1111	vi A	arg	_	1015	neu	rict	116		1020	urd	vsħ	ьсu	пуъ

Glu Ala Glu Val Ser Gln Glu Thr Val Leu Glu Thr Arg Trp Gln Glu 1030 1035 1040 Asn Val Leu Lys Phe Gln Pro Ser Thr Ser Lys Glu Ile Glu Asp Phe 1055 1050 1045 Leu Gln Ser Ala Pro Ser Ala Lys Ala Arg Lys Phe Ala Glu Lys His 1060 1065 1070 Leu Arg Thr Leu Ile Thr Gln Phe Asn Gly His Glu Arg Pro Pro Ser 1075 1080 1085 Glu Ala Thr Ala Val Pro Met Asp Tyr Thr Pro Thr Pro Ile Pro Thr 1090 1095 1100 Pro Gln Ala Val Ser Thr Ala Thr Ala Glu Lys Gly Lys Ala Ala Trp 1105 1110 1115 Asn Lys Ile Gln Gln Ala Phe Gln Asp Phe Asn Phe His Leu Ile Asp 1125 1130 1135 Ala Ser Asp Trp Gln Glu Met Ala Ser Glu Tyr Ser Arg His Gly Ser 1140 1145 1150 Ser Leu Pro Gly Thr Val Gly Pro Lys Leu Val Arg Phe Met Glu Ser 1160 1165 1155 Ile Ser Asn Thr Leu Asp Asp Ile Leu Thr Gln Lys Leu Ala Ser Leu 1170 1175 1180 Leu Pro Asn Gly Pro Ala Phe Arg Pro Pro Ala Phe Asp Trp Ile Ala 1185 1190 1195 1200 Pro Tyr Gln Thr Arg Val Asn Ala Phe Leu Lys Thr Ile Gly Leu Pro 1205 1210 Met Val Arg Asn Leu Ala Asp Lys Ile His His Gln Cys Gln Thr Val 1220 1225 1230 Ser His Ala Val Gln Ser Ala Asp Leu Gln Gln Ala Thr Val Gly Thr 1235 1240 1245 Ser Leu Glu Arg Pro Ala Ala Glu Tyr Cys Arg Ile Leu Ser Asp Met 1250 1255 1260 Gln Val Ala Phe Asn Asp His Gly Ile Ala Val Arg Ser Glu Ala Ala 1265 1270 1275 Ala Tyr Thr Asp Ala Ile Asn Ser Pro Ala Asn Val Val Thr Pro Pro 1285 1290 Lys Pro Asn Leu Glu Ala Pro Lys Lys Leu Ile Thr Ala Thr Asp Ala 1300 1305 1310 Leu Thr Val Glu Asp Phe Pro Asp Phe Leu Lys Thr Ser Ile Leu Gln 1315 1320 1325 Gln Glu Gln Arg Leu Ile Ala Leu Gln Arg Ala Glu Phe Gln Gln Leu 1335 1340 1330 Glu Ala Ser Ile Ser Ala Ala Glu Arg Leu Arg Gln Ser Thr Arg Asp 1345 1350 1355 1360 Glu Ile Ala Gly Lys Met Ala Thr Ala Ile Thr Gln Leu Leu Pro Arg 1365 1370 1375 Ala Pro Val Ala Ile Ser Ser Arg Pro Leu Asn Leu Ser Lys Pro Ile 1380 1385 1390 Asp Phe Leu Ser Ser Thr Val Tyr Asp Lys Ile Leu Asp Lys Glu Pro 1405 1395 1400 Tyr Glu Thr Ala Ile Ala Gly Phe Ala Trp Leu Glu Ile Ala Thr Lys 1410 1415 1420 Ser Val Met Val Tyr Ser Gln Gln Asn Glú Thr Gln Gln Leu Asn Val 1435 1430 Leu Leu Ser Glu Val Glu Lys Gln Ser Thr Val Ala Gln Arg Leu His 1445 1450 1455 Asp Leu Glu Leu Ser Ala Lys Asn Thr Asp Asp Val Lys Val Leu Lys 1460 1465 1470 Gln Ala Leu Asp Glu Leu Ala Pro Leu Arg Val Lys Gly Gly Lys Thr 1475 1480 1485 Thr Val Asp Ala Trp Lys Gln Lys Leu Glu Ser Ile Glu Ser Leu Leu 1500 1495 1490 Arg Ala Thr Arg Thr Ala Gly Glu Ile Ser Ser Glu Leu Glu Arg Ile 1505 1510 1515 Gly Thr Gln Ala Val Gly Thr Ile Thr Val Arg Asp Leu Gly Thr Leu 1525 1530 1535 Ser Asp Gln Cys Arg Glu Ala Ala Asn Phe Leu Arg Gln Ala Ser Leu 1540 1545 1550 Pro Glu Gly Phe Ser Asp Ile Gly Thr Lys Leu Ser Glu Leu Gln Ala 1565 1560 1555 Tyr Ile Lys Tyr Lys Lys Gln Phe Leu Glu His Phe Glu Thr Thr Gln 1570 1575 1580 Pro Asn Val Phe Gln Arg Phe Pro Leu Ser Gln Asn Ile Thr Glu Asn 1585 1590 1595 1600 Val Pro Ala Arg Pro Ala Met Asp Ser Val Ala Arg Leu Thr Asn His 1605 1610 1615 Leu His Val Arg Gly Ser Ala Pro His Phe Thr Thr Trp Ile Glu Thr 1620 1625 1630 Leu Pro Thr Val Asp Pro Glu Lys Pro Thr His Val Pro Ala His Gly 1635 1640 1645 Gly Ala Pro Leu His Arg Gln Ile Thr Tyr Ser Asn Val Leu Glu Ala 1650 1655 1660 Leu Phe Ser Leu Cys Ser Thr Thr Leu Thr Pro Val Pro Thr Ala Pro 1665 1670 1675 Gly Leu Glu Ile Ala Thr Arg Ala Arg Arg Gly Ala Glu Ala Ala Thr 1685 1690 1695 Trp Met Asp Arg Gln Trp Pro Asp Ile Ala Gln Thr Leu Gln Asp Val 1700 1705 1710 Leu Asp Thr Tyr Glu His Thr Thr Ala His Ala Asn Arg Asp Ala Ala 1715 1720 1725 Phe Asn Thr Phe Leu Ala Met Cys Val Phe Thr Gln Ile Ile Arg Gly 1740 1730 1735 Ala Ser Arg Ala Val Thr Leu Pro Lys Leu Pro Ser Thr Ala Val Asp 1745 1750 1755 1760 Phe Pro Glu Glu Ile Val Leu Thr Pro Arg Glu Cys Thr Thr Leu Val 1765 1770 1775 Thr Ala Met Trp Pro Thr Leu Ala Ala Ile Leu Arg Leu Lys Ser 1780 1785 1790 Tyr Ser Glu Ala Leu Gly Leu Met Ser Arg Phe Leu Pro Leu Met Phe 1795 1800 1805 Gln Ala Leu Pro His Leu Thr Leu Glu Ala Gln Val Lys Asn Gly Pro 1810 1815 1820 His Asn Thr Pro Pro Gln Leu Arg Cys Phe Ala Lys Thr Glu Ala Ile 1825 1830 1835 1840 Pro Tyr Phe Pro Ala Gln Trp Gln Ser Ala Asn Leu Glu Gln Ser Leu 1845 1850 1855 Trp Gly Gln Thr Asp Phe Leu Gln Ile Cys Asp Asn Asn Gln Arg Lys 1860 1865 1870 Ala Arg Val Ala Ala Val Thr Trp Ala Leu Thr Thr Ile Asp Gly Val 1875 1880 1885 Val Leu Asp Gln Leu Trp Ser Thr Phe Lys Pro Met Thr Ala Ala Ser 1895 1900 Asp Asp Thr Tyr Val Asp Leu Val Glu Thr Leu His Leu Thr Thr Phe 1905 1910 1915 Gly Pro Arg Gly Pro Thr Pro Arg Arg Glu Thr Thr Thr Glu His Pro 1925 1930 1935 Pro Tyr Glu Tyr Gly Gln Pro Thr Gly Tyr Cys Ile Ser Gly Gln Ser 1940 1945 1950 Thr Thr Pro Val Gln Ala Ser Asn Thr Pro Val Ser Ala Phe Glu Ala 1955 1960 1965 Val Leu Gly Ala Met Val Phe His Val Pro Ile Arg Ile Phe Leu Ala 1970 1975 1980 Ala Thr Pro Lys Arg Leu Gly Gln Ala Arg Gly Gly Met Gly Leu Leu 1995

Thr Pro Ile Leu Glu Cys Val Pro Asp Val Glu Pro Phe Lys Ser Leu 2005 2010 2015 Tyr Asn Ala Pro Arg Lys Pro Val Pro Ile Glu Thr Leu Pro Ala Ser 2020 2025 2030 Leu His Pro His Asp Glu Arg Gln Val Phe Leu Arg Gln Ala Gln Trp 2045 2035 2040 Leu Ser Tyr Arg Phe Thr Pro His Glu Ala Ala Arg Ser Ser Thr Pro 2050 2055 2060 Pro Leu Leu Val Val Ile Asp Pro Glu Asn Leu Val Thr Ala Thr Tyr 2065 2070 2075 2080 Ser Ser Gly Gly Pro Ala Asn Phe Glu Ser Arg Pro Phe Tyr Val Met 2085 2090 2095 Pro Gly Pro Tyr Pro Pro Asp Trp Pro Lys Thr Leu Ser Val Thr Ser 2100 2105 2110 Asn Thr Ser Val Thr His Leu Ser His Asp Glu Ile Cys Asn Leu Phe 2115 2120 2125 Thr Thr Leu Ser Arg Glu His Gly Thr Val Gln Gly Arg Asp Ile Phe 2140 2130 2135 Ala Ala Ala Pro Thr Asn Val Thr Pro Glu Gln Thr Ala Asn Pro Pro 2145 2150 2155 2160 Ala Trp Glu Thr Asp Asn Arg Leu Ile Thr Gln Thr Glu Thr Ala Lys 2165 2170 2175 Lys Pro His Ile Ile Pro Ala Ser Pro Lys Ala Arg Thr Asp Pro Pro 2180 2185 2190 Val Glu Thr Thr His His Ser Gln Gly Gln Ala Ser Gln His Ala 2205 2195 2200 Asn Ser Asn Val Asn Gln Pro Gly Gln Ile Thr Ser His Ala Ser Arg 2220 2215 2210 Asn Thr Pro Ser Thr Ala Pro Gln Ala Ser Ser Ser Pro Glu Lys Phe 2225 2230 2235 2240 Asn Thr Gln Thr Val Pro Arg Leu Ile Ser Gln Thr Ser Glu Thr Ala 2245 2250 2255 His Ile Asn Gln Pro Ala Ser Gly Gln Val Thr Glu Pro Lys Gly Ile 2260 2265 2270 Phe Gly Thr Tyr Lys Pro Arg Val Leu Thr Glu Pro Ala Lys Pro Ala 2275 2280 2285 Asn Ala Gly Val Ala Ser Arg Gln Pro Glu Ala Thr Thr Thr Val Pro 2290 2295 2300 Lys Leu Pro Ile Asn Pro Pro Thr Ala Arg Val Phe Ile Gly Thr Ala 2305 2310 2315 2320 Ser Lys Leu Ser Pro Ala Val Glu Glu Ser His Gly Ala Thr Pro Asp 2325 2330 2335 Ala His Gln Ser Lys Ile Asp Arg Glu Lys Tyr Ala Glu Ser Arg Pro 2340 2345 2350 Arg Arg Thr Pro His Leu Glu Glu Gly Pro Arg Glu Pro His Val Asn 2360 2365 Thr Pro Thr Ser Ala His Ile Asn Val Pro Ser Ser Gln Gly Gln Lys 2370 2375 2380 Thr Val His Gly Arg Glu Asn Pro Gly Leu Gln Thr Ala Thr Pro Ser 2385 2390 2395 Ala Pro Gln Pro Thr Ala Ser Asn Pro Arg Ile Gln Tyr Thr Leu Pro 2405 2410 2415 Arg Thr Asp Gly Arg Leu Leu His Asp Glu Ser Glu Val Glu Ser Thr 2420 2425 2430 Pro Thr Glu Glu Val Lys Arg Ser Pro Lys Thr Gln Asp Val Ser His 2435 2440 2445 Gly Pro Glu Pro Asp Asp Ser Arg Trp Thr Ala Pro Leu Gly Pro Thr 2450 2455 2460 Ile Glu Ile His Arg Leu Glu His Pro Gln Ile Leu Lys Asn Ile Thr 2470 2475 Ser Leu Thr Val Pro Thr Pro Arg Val Thr Pro Ile Pro Pro Thr Asn

2495 2490 2485 Ile Trp Ile Pro Leu Ser His Val Asn Ile Gln His Glu Glu Ile Thr 2500 2505 2510 Arg Ala Lys Asn Val Leu Met Arg Phe Ile Gln Asn Val Arg Arg Lys 2525 2515 2520 Leu Gln Ala Ser Ser Asp Ala Leu Ser Glu Ala Ile Ala Arg Ile Lys 2535 Phe Leu Tyr Leu 2545 <210> 144 <211> 510 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(510) <400> 144 atg teg tee ttg egg gtt aag gag eea ate gtt eag gga ege ett gag Met Ser Ser Leu Arg Val Lys Glu Pro Ile Val Gln Gly Arg Leu Glu 5 10 cac gat tac cca aat cac ccg ctg gtt gcc gag atg aac aac ctt ccc His Asp Tyr Pro Asn His Pro Leu Val Ala Glu Met Asn Asn Leu Pro cag ggt gac atg tct ccc gcc cag tat gcc atc gcg aaa cgt aat tat 144 Gln Gly Asp Met Ser Pro Ala Gln Tyr Ala Ile Ala Lys Arg Asn Tyr 35 40 ctg gtg ttt tta acg gcc aaa cat cat tac gac atg tat atg caa aag Leu Val Phe Leu Thr Ala Lys His His Tyr Asp Met Tyr Met Gln Lys 50 aag aat gga att ctg_cgc aaa gac cac ctc cgg ggc ctt cgc. ggc aaa 240 Lys Asn Gly Ile Leu Arg Lys Asp His Leu Arg Gly Leu Arg Gly Lys aag gac get agt tet agt ate teg gge gtt ttg tee ggg tee gge teg Lys Asp Ala Ser Ser Ser Ile Ser Gly Val Leu Ser Gly Ser Gly Ser 90 ged ged deg age gtt ged deg gtg ged ted add etd ggg tea aat age 336 Ala Ala Pro Ser Val Ala Pro Val Ala Ser Thr Leu Gly Ser Asn Ser 105 ttt act acg atc tcc agc ggg ccc cat tca ttg ata ggc tcg atg ggc 384 Phe Thr Thr Ile Ser Ser Gly Pro His Ser Leu Ile Gly Ser Met Gly 120 115 ecc geg eec ggt gge gge gga eec ggg age gtg geg tet tet gge ata Pro Ala Pro Gly Gly Gly Pro Gly Ser Val Ala Ser Ser Gly Ile 140 135 130 ggg tee aet teg tta tea eet age gae gea aet aet etg gae aea aga 480 Gly Ser Thr Ser Leu Ser Pro Ser Asp Ala Thr Thr Leu Asp Thr Arg 155 150 145

510 cgc tcg tct caa aat aaa aaa agc aag tga Arg Ser Ser Gln Asn Lys Lys Ser Lys 165 <210> 145 <211> 169 <212> PRT <213> Macaca mulatta rhadinovirus 17577 Met Ser Ser Leu Arg Val Lys Glu Pro Ile Val Gln Gly Arg Leu Glu 10 His Asp Tyr Pro Asn His Pro Leu Val Ala Glu Met Asn Asn Leu Pro 30 . 20 25 Gln Gly Asp Met Ser Pro Ala Gln Tyr Ala Ile Ala Lys Arg Asn Tyr 4.0 35 Leu Val Phe Leu Thr Ala Lys His His Tyr Asp Met Tyr Met Gln Lys 60 55 Lys Asn Gly Ile Leu Arg Lys Asp His Leu Arg Gly Leu Arg Gly Lys 70 Lys Asp Ala Ser Ser Ser Ile Ser Gly Val Leu Ser Gly Ser Gly Ser 95 90 85 Ala Ala Pro Ser Val Ala Pro Val Ala Ser Thr Leu Gly Ser Asn Ser 100 105 110 Phe Thr Thr Ile Ser Ser Gly Pro His Ser Leu Ile Gly Ser Met Gly 120 125 Pro Ala Pro Gly Gly Gly Bro Gly Ser Val Ala Ser Ser Gly Ile 140 135 Gly Ser Thr Ser Leu Ser Pro Ser Asp Ala Thr Thr Leu Asp Thr Arg 150 155 Arg Ser Ser Gln Asn Lys Lys Ser Lys 165 <210> 146 <211> 1347 <212> DNA <213> Macaca mulatta rhadinovirus 17577 <220> <221> CDS <222> (1)..(1347) atg gcc tcg ggc cgc ttg cct aac ctg gct gaa gac gaa gct gcc tgt Met Ala Ser Gly Arg Leu Pro Asn Leu Ala Glu Asp Glu Ala Ala Cys 15 5 10 cat ggg cgc ggt tot tat cct gcc cat cgt tgg ctg gat ggt tot cgg 96 His Gly Arg Gly Ser Tyr Pro Ala His Arg Trp Leu Asp Gly Ser Arg 2.0 25 ctg ggc tta gat ctc gcg gcc tct ata cgc tca atc gga cta tgc ccc Leu Gly Leu Asp Leu Ala Ala Ser Ile Arg Ser Ile Gly Leu Cys Pro

192

40

gaa tgc tac gtg tgt ttt gtg acg tac ggg ctc ggt gcc tgg gac gga

Glu Cys Tyr Val Cys Phe Val Thr Tyr Gly Leu Gly Ala Trp Asp Gly

	50					55					60				
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											_	-	aac Asn		288
													cgg Arg 110		336
													ggc Gly		384
													agg Arg		432
													tgt Cys		480
													acc Thr		528
				_	_	-		_	_		_		tgg Trp 190	_	576
													tcc Ser		624
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_													ctg Leu		768
_		_	_		_					_			acg Thr 270		816
													ctg Leu		864
													agc Ser		912

gag Glu 305	aag Lys	acc Thr	gtg Val	cca Pro	gct Ala 310	ccg Pro	agg Arg	gtg Val	gtg Val	gtg Val 315	tgc Cys	ctc Leu	gag Glu	tgc Cys	ggt Gly 320	960
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ccc Pro	acc Thr	aac Asn	gtg Val 340	ttt Phe	ttc Phe	agt Ser	cgc Arg	gac Asp 345	caa Gln	aaa Lys	gag Glu	aag Lys	cag Gln 350	ctt Leu	tcg Ser	1056
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tat Tyr 385	tta Leu	cgg Arg	tgc Cys	gtg Val	ctc Leu 390	gcc Ala	aac Asn	aac Asn	gcg Ala	gca Ala 395	cat His	gcc Ala	ata Ile	cga Arg	gac Asp 400	1200
gcg Ala	aac Asn	tcc Ser	ctg Leu	gtt Val 405	agc Ser	gtc Val	gtc Val	gtg Val	ccc Pro 410	tgt Cys	ttg Leu	gcg Ala	tcg Ser	ccg Pro 415	gac Asp	1248
tgc Cys	gcg Ala	acc Thr	ggc Gly 420	cta Leu	tta Leu	aag Lys	cat His	ttg Leu 425	cgt Arg	gtg Val	gcc Ala	gag Glu	ctg Leu 430	ttt Phe	tat Tyr	1296
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taa														• -		1347

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<213> Macaca mulatta rhadinovirus 17577

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110

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Leu Glu Ala Tyr Ala Trp Val Leu Arg Cys Ile Cys Thr Gly Val Gly
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                                             125
Cys Pro Ser Asp Glu Gly Leu Ser Leu Thr Ala Val Pro Arg Ser Ala
                                      140
                      135
Trp Ser Arg Tyr Leu Val Val Ser Phe Gln Arg Ala Cys Cys Leu Val
                                      155
                150
Cys Lys Thr Leu Asn Cys Arg Gln Arg Phe Pro Leu Val Thr Cys Leu
               165
                                  170
Pro Gln His Ala Leu Asp Leu Pro Val Leu Arg Lys Lys Trp Asn Gly
                             185
                                                 190
Gly Gly Cys Val Ser Met Gln Leu Asn Val Pro Ser Ile Ser Arg Arg
                          200
                                             205
       195
Leu Gly Ala Asn Leu Asn Glu Ser Val Pro Gly Pro Ser Asp Ala Gly
                                         220
            215
Leu Leu Ala Ser Leu Arg Glu Leu Ala Pro Thr Val Pro Cys Gly Asn
                230
                          235
Pro Phe Asn Ala; Leu Leu Arg Ser Leu Thr Phe Arg Ala Leu Leu Ser
                                  250
               245
Met Ser Arg Val Val Leu Pro Ile Gly Glu Ser Thr Glu Thr Glu Ile
           260
                              265
Ser Arg Asp Leu Gly Gln Lys Val Leu Ala Tyr Asn Val Leu Phe Pro
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                                             285
Cys Ile Ser Leu Pro Val Trp Ser Gln Val Val Ala Arg Ser Val Leu
                                          300
                      295
Glu Lys Thr Val Pro Ala Pro Arg Val Val Cys Leu Glu Cys Gly
                310
                                     315
Tyr Cys Leu Asn Phe Gly Arg Gly Lys Phe Glu Thr Val Asn Phe Pro
              325
                                  330
Pro Thr Asn Val Phe Phe Ser Arg Asp Gln Lys Glu Lys Gln Leu Ser
                              345
Ile Cys Ala Thr Thr Gly Arg Val Tyr Cys Ser Tyr Cys Gly Gly Ser
                                             365
                         360
His Met Arg Val Ile Ser Leu Phe Glu Ile Thr Cys Val Gly Asp Pro
                      375
                                          380
Tyr Leu Arg Cys Val Leu Ala Asn Asn Ala Ala His Ala Ile Arg Asp
               390
                                   395
Ala Asn Ser Leu Val Ser Val Val Pro Cys Leu Ala Ser Pro Asp
              405
                                 410
                                                 415
Cys Ala Thr Gly Leu Leu Lys His Leu Arg Val Ala Glu Leu Phe Tyr
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Leu Thr Ser Ser Ile Ser Ser Leu Ser Cys Gly Lys Cys Asn Arg Ser
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tct tac ctg gga ccc tct gga atc tcc ctg gac ttg gag agg tgt cag
Ser Tyr Leu Gly Pro Ser Gly Ile Ser Leu Asp Leu Glu Arg Cys Gln
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30 2.5 144 gac ggg gct ccc gta tac gct aaa ggc ggg gcg gtc ccc gtg tgc acc Asp Gly Ala Pro Val Tyr Ala Lys Gly Gly Ala Val Pro Val Cys Thr 4.0 3.5 gtg cgc ctg cag cac ggc tgc gtc tat cat ctc gag ttt gtg tat aag Val Arg Leu Gln His Gly Cys Val Tyr His Leu Glu Phe Val Tyr Lys ttt tgg ctc cac aaa cta gag aga ctg gcc tac ccg ttt gcc ccg tgt 240 Phe Trp Leu His Lys Leu Glu Arg Leu Ala Tyr Pro Phe Ala Pro Cys 65 288 tit gta att atc aac aac ggt tig gcc acc acg cig aaa igt tit tig Phe Val Ile Ile Asn Asn Gly Leu Ala Thr Thr Leu Lys Cys Phe Leu 90 85 tgt aag cca cgt gac gcc gat gcc cag ttt gga aaa aac ctg cct ata 336 Cys Lys Pro Arg Asp Ala Asp Ala Gln Phe Gly Lys Asn Leu Pro Ile 105 aat tog gao gtg tat ott gag agg aac tog too gtg too otg ggo cag 384 Asn Ser Asp Val Tyr Leu Glu Arg Asn Ser Ser Val Ser Leu Gly Gln 120 gac gat ttt atg aaa ttt aag gca cgt ctg gtt ttc tcc gga gac cta Asp Asp Phe Met Lys Phe Lys Ala Arg Leu Val Phe Ser Gly Asp Leu 135 140 130 aac gtt tac agc tcc atg gtc ata tgc cgc acc tac ttt acg gag cac 480 Asn Val Tyr Ser Ser Met Val Ile Cys Arg Thr Tyr Phe Thr Glu His 150 528 cga cag gtt tta cag ttt ttg gtc gtg act cca aag agc gct aaa cgg Arg Gln Val Leu Gln Phe Leu Val Val Thr Pro Lys Ser Ala Lys Arg 165 170 tta aaa acc ctt ctt aga acg gtt ttt gcc ctg acg ggt cac tcc gac 576 Leu Lys Thr Leu Leu Arg Thr Val Phe Ala Leu Thr Gly His Ser Asp 185 624 ggc ctc ggt gcg ttg agg cga acg ggc tec gtg gcc cgc cct teg ggg Gly Leu Gly Ala Leu Arg Arg Thr Gly Ser Val Ala Arg Pro Ser Gly tcg gag ttg aag gat att ggg cgc gga gag cgt gcg gcg atg acc aat 672 Ser Glu Leu Lys Asp Ile Gly Arg Gly Glu Arg Ala Ala Met Thr Asn 215 220 675 taa 225 <210> 149

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acc Thr	tgc Cys	cta Leu	tac Tyr	gcc Ala 85	ccc Pro	cga Arg	tca Ser	tgg Trp	acg Thr 90	gct Ala	acc Thr	ctc Leu	atg Met	gtg Val 95	gct Ala	288
gcc Ala	gac Asp	ctt Leu	ttg Leu 100	gaa Glu	cta Leu	acg Thr	cac His	gtg Val 105	tac Tyr	ttc Phe	ccg Pro	caa Gln	tgc Cys 110	gtg Val	aaa Lys	336
gat Asp	Gly ggg	cca Pro 115	gta Val	tac Tyr	acc Thr	gcc Ala	caa Gln 120	agc Ser	atc Ile	ctc Leu	gga Gly	atc Ile 125	gac Asp	gtc Val	cag Gln	384
ctg Leu	cac His 130	ttc Phe	ttc Phe	gca Ala	acc Thr	cgc Arg 135	tgc Cys	ttc Phe	cga Arg	ccc Pro	atc Ile 140	gac Asp	aga Arg	gaa Glu	caa Gln	432
ata Ile 145	ctc Leu	cac His	aca Thr	tct Ser	cat His 150	tta Leu	aat Asn	ttt Phe	tta Leu	caa Gln 155	acc Thr	gag Glu	ttt Phe	att Ile	agg Arg 160	480
ggc Gly	atg Met	tta Leu	gaa Glu	ggc Gly 165	acg Thr	att Ile	ccg Pro	gga Gly	tcg Ser 170	ttc Phe	tgt Cys	ttt Phe	aaa Lys	acg Thr 175	tcc Ser	528
tgg Trp	ccg Pro	cgc Arg	aca Thr 180	gaa Glu	aag Lys	gac Asp	gac Asp	caa Gln 185	caa Gln	cct Pro	acc Thr	gtt Val	gcg Ala 190	tgt Cys	tgt Cys	576
Ser	gtt Val	Gly 195	Arg	Gly	Ser	His	Thr 200	Asn	Arg	Asp	Asn	Arg 205	Leu	Pro	Glu	624
Āsp	ctg Leu 210	Glu	Glu	Ala	Phe	Asn 215	Ser	Thr	Asn	Ala	Glu 220	Glu	Lys	Pro	Ser	672
Leu 225		Gly	Val	Phe	Ser 230	Ala	Thr	Trp	Ala	Glu 235	Ser	Gln	Leu	Leu	Gly 240	720
Ser	gac Asp	Thr	Gln	Gln 245	Ala	Asp	Thr	His	Leu 250	Gln	Pro	Ser	Ala	Phe 255	Pro	768
Thr	Pro	Glu	Asp 260	Ala	Asp	Gln	Ser	Gln 265	Gly	Pro	Cys	Leu	Met 270	His		816
Thr	Leu	Asn 275	Leu	Lys	Thr	Lys	Asn 280	His	Thr	Ala	Ser	1le 285	Cys	Val	cta Leu	864
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aat	. cgc	ata	tcg	tac	atc	cta	aac	gat	ccg	gac	tca	ctç	l tca	cac	gtg	1008

Asn	Arg	Ile	Ser	Tyr 325	Ile	Leu	Asn	Asp	Pro 330	Asp	Ser	Leu	Ser	His 335	Val	
										cgg Arg						1056
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tcg Ser	cac His 370	tgt Cys	ccc Pro	gcg Ala	gtt Val	tta Leu 375	ttt Phe	aaa Lys	tgc Cys	cca Pro	cct Pro 380	ccc Pro	gaa Glu	aag Lys	tat Tyr	1152
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aga Arg	ata Ile	ttt Phe	gac Asp	tgc Cys 405	gag Glu	acc Thr	tta Leu	cag Gln	acc Thr 410	ctg Leu	gcc Ala	gtc Val	ctc Leu	ttt Phe 415	aag Lys	1248
Gly aaa	tct Ser	caa Gln	ctg Leu 420	gcc Ala	aaa Lys	atc Ile	ggc Gly	aaa Lys 425	acc Thr	acg Thr	tcg Ser	ctc Leu	gag Glu 430	ata Ile	atc Ile	1296 ,
cgt Arg	gaa Glu	ctc Leu 435	gga Gly	ttt Phe	caa Gln	ctg Leu	cgt Arg 440	cga Arg	cac His	aac Asn	att Ile	caa Gln 445	atc Ile	acc Thr	cac His	1344
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1				5					10	Met				15		
		_	20					25		Pro			30			
Ala	Phe	Asn 35	Ser	Pro	Val	Leu	Ile 40	His	Thr	Gln	Asp	Ser 45	Leu	Gln	Pro	
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	Cys	Leu	Tyr	Ala 85		Arg	Ser	Trp	Thr 90	Ala	Thr	Leu	Met	Val 95		
Ala	Asp	Leu			Leu	Thr	His			Phe	Pro	Gln	Cys 110		Lys	
Asp	Gly		100 Val	Tyr	Thr	Ala		105 Ser	Ile	Leu	Gly			Val	Gln	
Leu		115 Phe	Phe	Ala	Thr		120 Cys	Phe	Arg	Pro		125 Asp	Arg	Glu	Gln	
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Trp Pro Arg Thr Glu Lys Asp Asp Gln Gln Pro Thr Val Ala Cys Cys
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Asp Leu Glu Glu Ala Phe Asn Ser Thr Asn Ala Glu Glu Lys Pro Ser
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Leu Leu Gly Val Phe Ser Ala Thr Trp Ala Glu Ser Gln Leu Leu Gly
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Ser Asp Thr Gln Gln Ala Asp Thr His Leu Gln Pro Ser Ala Phe Pro
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Thr Pro Glu Asp Ala Asp Gln Ser Gln Gly Pro Cys Leu Met His Pro
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                                                 270
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Thr Leu Asn Leu Lys Thr Lys Asn His Thr Ala Ser Ile Cys Val Leu
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Cys Glu Cys Leu Ala Ala His Pro Asp Ala Gly Pro Val Leu Lys Asp
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Leu Arg Arg Asp Ile Leu Glu Asn Met Glu Asn Asn Val Lys Leu Val
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                310
Asn Arg Ile Ser Tyr Ile Leu Asn Asp Pro Asp Ser Leu Ser His Val
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               325
Arg Asp Glu His Leu Arg Gly Leu Ile Lys Arg Cys Ser Ala Gln Glu
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                                                 350
Ile His Lys His Phe Phe Cys Asp Pro Val Cys Val Leu Asn Thr Tyr
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Ser His Cys Pro Ala Val Leu Phe Lys Cys Pro Pro Pro Glu Lys Tyr
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Lys Lys Leu Lys Ala Arg Leu Ala Thr Gly Glu Phe Leu Asp Cys Asn
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Arg Ile Phe Asp Cys Glu Thr Leu Gln Thr Leu Ala Val Leu Phe Lys
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Gly Ser Gln Leu Ala Lys Ile Gly Lys Thr Thr Ser Leu Glu Ile Ile
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                                                                96
Pro Ser Val Arg Arg Pro Asp Gly Pro Gln Ser Thr Arg Pro Ala Ser
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gtg Val 65	gac Asp	ttt Phe	tta Leu	aga Arg	gaa Glu 70	atg Met	999 Gly	acc Thr	ccg Pro	ata Ile 75	tgc Cys	acc Thr	tca Ser	aag Lys	tcc Ser 80	240
gtt Val	atg Met	ttg Leu	ccg Pro	tta Leu 85	aac Asn	cta Leu	aaa Lys	acc Thr	atc Ile 90	gcc Ala	ccg Pro	ggt Gly	cgg Arg	tgc Cys 95	gtc Val	288
tct Ser	ctc Leu	tca Ser	tca Ser 100	ttc Phe	gga Gly	cac His	tcg Ser	tca Ser 105	aac Asn	atg Met	Gly 333	ttc Phe	aac Asn 110	tgt Cys	tcg Ser	336
tcg Ser	tgc Cys	acg Thr 115	cca Pro	act Thr	gac Asp	agg Arg	tca Ser 120	gcg Ala	gtg Val	tct Ser	ctg Leu	gac Asp 125	gca Ala	aac Asn	gcg Ala	384
ctc Leu	ggc Gly 130	gaa Glu	gat Asp	tcc Ser	gcc Ala	agg Arg 135	aaa Lys	aac Asn	agc Ser	gag Glu	ctg Leu 140	tgt Cys	tca Ser	gtg Val	gcg Ala	432
tta Leu 145	acc Thr	ttt Phe	tac Tyr	cac His	cac His 150	gcc Ala	gaa Glu	aag Lys	gtc Val	gtg Val 155	cag Gln	cac His	aag Lys	ggc Gly	ttt Phe 160	480
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cac His	gat Asp	cct Pro 195	tta Leu	cct	att Ile	ttt Phe	aca Thr 200	gtc Val	gat Asp	gcc Ala	gat Asp	gag Glu 205	aga Arg	ctc Leu	gca Ala	624
ctc Leu	tgg Trp 210	gcg Ala	gtg Val	ttc Phe	cac His	act Thr 215	aga Arg	gac Asp	cta Leu	cac His	ctg Leu 220	ggg Gly	gaa Glu	acc Thr	agt Ser	672
ctg Leu 225	Arg	ctc Leu	att Ile	atg Met	gac Asp 230	aac Asn	ctt Leu	cca Pro	aat Asn	tat Tyr 235	gac Asp	ata Ile	acg Thr	gtg Val	gac Asp 240	720
tgc Cys	atc Ile	aag Lys	caa Gln	acg Thr 245	tac Tyr	ata Ile	atg Met	aag Lys	ttt Phe 250	aca Thr	ccc Pro	tcg Ser	cga Arg	ceg Pro 255	gac Asp	768
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acc Thr	cta Leu	gac Asp	tgc Cys	acc Thr	gac Asp	gag Glu	ttt Phe	cga Arg	gaa Glu	gaa Glu	att Ile	caa Gln	agg Arg	ggc Gly	acg Thr	864

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275 280

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    3.5
Ser Asp Thr Gln Phe Phe Ser Ala Leu Thr Arg Arg His Glu Leu Gly
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Val Asp Phe Leu Arg Glu Met Gly Thr Pro Ile Cys Thr Ser Lys Ser
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Val Met Leu Pro Leu Asn Leu Lys Thr Ile Ala Pro Gly Arg Cys Val
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Ser Leu Ser Ser Phe Gly His Ser Ser Asn Met Gly Phe Asn Cys Ser
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                          105
          100
Ser Cys Thr Pro Thr Asp Arg Ser Ala Val Ser Leu Asp Ala Asn Ala
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                                       125
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Leu Gly Glu Asp Ser Ala Arg Lys Asn Ser Glu Leu Cys Ser Val Ala
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Leu Thr Phe Tyr His His Ala Glu Lys Val Val Gln His Lys Gly Phe
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Tyr Leu Ser Leu Leu Ser His Ser Met Glu Val Val Arg Lys Ser Phe
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His Asp Pro Leu Pro Ile Phe Thr Val Asp Ala Asp Glu Arg Leu Ala
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Leu Trp Ala Val Phe His Thr Arg Asp Leu His Leu Gly Glu Thr Ser
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Leu Arg Leu Ile Met Asp Asn Leu Pro Asn Tyr Asp Ile Thr Val Asp
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                         235
Cys Ile Lys Gln Thr Tyr Ile Met Lys Phe Thr Pro Ser Arg Pro Asp
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                              250
            245
Asn Ala Thr Val Thr Val Pro Val Asn Ser Ile Cys Glu Ala Val Ala
        260 265
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gac Asp	gac Asp	aga Arg	gag Glu 20	gcc Ala	gtg Val	ctg Leu	tgg Trp	ctt Leu 25	ttt Phe	gac Asp	cgg Arg	ccg Pro	gcg Ala 30	tcc Ser	gac Asp	96
gat Asp	acg Thr	ccc Pro 35	gag Glu	999 Gly	ttt Phe	gca Ala	aac Asn 40	999 Gly	ctg Leu	tgc Cys	ccc Pro	tca Ser 45	act Thr	gga Gly	gaa Glu	144
ccc Pro	ggt Gly 50	att Ile	ccc Pro	ctc Leu	ccg Prọ	gtg Val 55	ttg Leu	ctg Leu	gag Glu	gcc Ala	gtg Val 60	ttt Phe	ctc Leu	gtt Val	Gly aaa	192
cga Arg 65	ttg Leu	gac Asp	ctg Leu	gtc Val	tcc Ser 70	acc Thr	ttt Phe	ttt Phe	tta Leu	cta Leu 75	gac Asp	gtg Val	gga Gly	ttt Phe	att Ile 80	240
atc Ile	gag Glu	agg Arg	ctc Leu	cgg Arg 85	tcc Ser	agc Ser	ccc Pro	agt Ser	tac Tyr 90	ttt Phe	agt Ser	cca Pro	tac Tyr	aaa Lys 95	cac His	288
ctg Leu	atg Met	ctc Leu	tcc Ser 100	att Ile	gac Asp	cgc Arg	cag Gln	ctc Leu 105	tca Ser	gaa Glu	agg Arg	gac Asp	gtg Val 110	aaa Lys	aat Asn	336
tta Leu	gtt Val	ttt Phe 115	cta Leu	acg Thr	ggc	gac Asp	cag Gln 120	ctt Leu	ggt Gly	cgc Arg	agg Arg	cgc Arg 125	aac Asn	cag Gln	tca Ser	384
ccc Pro	acc Thr 130	ttt Phe	ttt Phe	cgg Arg	tgg Trp	ctc Leu 135	tcg Ser	caa Gln	atg Met	gaa Glu	aag Lys 140	gcg Ala	gcc Ala	ctg Leu	gtc Val	432
agc Ser 145	ccc Pro	tca Ser	aac Asn	tac Tyr	atg Met 150	gtt Val	tta Leu	agt Ser	gac Asp	ctg Leu 155	ctg Leu	cag Gln	gcc Ala	gtc Val	tcc Ser 160	480
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<213> Macaca mulatta rhadinovirus 17577

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gcg gat tta ctg aaa cag gaa aaa tca atc ctt aag gct tta agg tgg

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gte Va	g gag l Glu	tct Ser	gtt Val 180	cac His	aaa Lys	gcc Ala	atc Ile	gtg Val 185	aac Asn	ccg Pro	gcc Ala	acc Thr	ggc Gly 190	ggt Gly	ctg Leu	576
Pro	e eeg o Pro	tcc Ser 195	ctg Leu	gtg Val	gcg Ala	gcc Ala	gcc Ala 200	tgc Cys	gcg Ala	ctg Leu	ttt Phe	agc Ser 205	ctc Leu	ggt Gly	gcc Ala	624
gc	gcg a Ala 210	ccg Pro	cct Pro	ccg Pro	gcc Ala	aga Arg 215	ttg Leu	gcg Ala	gag Glu	gcc Ala	gtc Val 220	ggc Gly	gtt Val	tcg Ser	gcc Ala	672
gca Ala 22!	a Thr	ctc Leu	gcg Ala	gcc Ala	gcc Ala 230	gcc Ala	gag Glu	tcg Ser	gtt Val	gcc Ala 235	acc Thr	acc Thr	ttg Leu	cgg Arg	gaa Glu 240	720
tt! Phe	e Asp	gaa Glu	gac Asp	cac His 245	att Ile	tta Leu	agt Ser	aac Asn	gcc Ala 250	cgc Arg	ggt Gly	tcg Ser	tcg Ser	tga 255		765
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Ala Thr Leu Ala Ala Ala Ala Ala Glu Ser Val Ala Thr Thr Leu Arg Glu 225

Phe Asp Glu Asp His Ile Leu Ser Asn Ala Arg Gly Ser Ser 245

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	_	cag Gln		_			_			_						528
_	_	cct Pro			_	-										576
		caa Gln 195	-		_	_	_		-							624
ccc Pro	gac Asp 210	tca Ser	ccg Pro	gga Gly	cca Pro	ccc Pro 215	caa Gln	tcg Ser	cca Pro	acg Thr	cct Pro 220	caa Gln	cag Gln	gcc Ala	cca Pro	672
		aac Asn	_		_		-	-			_				_	720
		cgg Arg				_										768
		gjà aaa														816
		ctc Leu 275														864
		caa Gln														912
		ccg Pro														960
		tca Ser														1008
_		tat Tyr	_	_	-	_										1056
		tgg Trp 355														1104
		cgc Arg														1152
		cgc Arg														1200

ttt Phe	tgc Cys	att Ile	acg Thr	gtg Val 405	Phe	tgt Cys	caa Gln	agc Ser	cgc Arg 410	gga Gly	acc Thr	gcc Ala	aag Lys	gcc Ala 415	gtc Val	1248
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			Ser				atg Met 440									1344
taa																1347
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	Arg	Ser	Ser 20		Arg	Gly	His	Cys 25		Arg	Arg	Gly	Gly 30		Arg	
Glu	Gln	Ala 35	Gly	Arg	Arg	Gly	Arg 40	Gly	Arg	Gly	Thr	Ala 45		Pro	Ala	
Ala	Ala 50	Pro	Ala	Pro	Pro	Ala 55	Pro	Thr	Thr	Ser	Gly 60		Gln	Val	Arg	
Ala 65	Val	Ala	Glu	Gln	Gly 70		Gly	Ser	Asp	Thr 75	Glu	Thr	Ala	Thr	Glu 80	
Ser	Arg	His	Gly	Ser 85	Ser	Gln	Gly	Ser	Pro 90	Ser	Gly	Ser	Gly	Ser 95	Glu	
Ser	Val	Ile	Val 100	Leu	Gly	Ser	Pro	Thr 105	Pro	Ser	Pro	Ser	Gly 110	Ser	Ala	
		115					Ser 120					125				
	130					135	Ser				140					
45					150		Pro			155					160	
				165			Leu		170					175		
lu	Pro	Pro	Glu 180	Pro	Pro	Thr	Ser	Leu 185	Pro	Pro	Pro	Asp	Ser 190	Pro	Gly	
		195					Thr 200					205				
	210					215	Gln				220					
Ser 225	Pro	Asn	Thr	Gln	Gln 230	Ala	Val	Ser	His	Thr 235	Asp	His	Pro	Thr	Gly 240	
?ro	Ser	Arg	Pro	Gly 245	Pro	Pro	Phe	Pro	Gly 250	His	Thr	Ser	His	Ser 255	Tyr	
ſhr	Val	Gly	Gly 260	Trp	Gly	Pro	Pro	Thr 265	Arg	Ala	Gly	Gly	Val 270		Cys	
eu	Arg	Leu 275	Arg	Cys	Thr	Ser	His 280	Asn	Ser	His	Glu	Asp 285		Ala	Pro	
~ T	7	a1	015	Q1	<b>a</b> 1-	<b>a</b> 1	01	<b>~</b> 1.	~ 7	_	~ 7	~ 7	~ 3	_	~ 7	

300

Glu Arg Gln Glu Glu Glu Glu Glu Glu Glu Gln Gln Pro Ala

Arg Pro Pro Arg Pro Pro Arg Pro Pro Arg Tyr Pro Ile Pro

305	310		315		320
Tyr Pro Ser Se	er Glu Glu 325	Glu Val Pi	ro Arg Lys 330	Tyr Arg Pr	o Gln Arg 335
Arg Phe Tyr Ai	_	-	ro Arg Ile 45	Asp Pro Pr 35	_
Gly Pro Trp Cy 355	s His Gly	Val Ile Pl 360	he Cys Asn	Ser Asp Pr 365	o Tyr Ser
Leu Tyr Arg Lo	eu Ala Arg	Cys Leu G	ln Phe Pro	Gly Ile Ar	g Ala Ser
Ser Val Arg Va 385	al Leu Pro 390	Asp Ala Pr	ro Gly Ser 395	Pro Val Il	e Pro Ala 400
Phe Cys Ile Th	nr Val Phe 405	Cys Gln Se	er Arg Gly 410	Thr Ala Ly	s Ala Val 415
Lys Lys Ala Ai 42	3 3	-	rg His His 25	Pro Ser Al	
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Met Ser Gly Gl		_			
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Val Arg Cys Al		Thr His Ty		Val Pro Va	
acc gcg tcc ct	g ggg tgc	gtg tta a	cg aca ccc	cac gac gt	t ctt atc 144
Thr Ala Ser Le	u Gly Cys	Val Leu Ti 40	hr Thr Pro	His Asp Va 45	l Leu Ile
gtt acc tgg ca	_	_	_	_	
Val Thr Trp Gl 50	n Lys Gin	55	ro Ser Pro	60	I Aia Thr
tat agt tcc ga Tyr Ser Ser Gl					
65	70	III VAI V	75	PLO PILE AI	80
gtt gac att co Val Asp Ile Pr	-	_ (ii)		_	7
var noh ite er	85	Lyo neu II	90	III Dea by	95
aat gcc acc ct			_	_	
Asn Ala Thr Le	_		os Tyr Leu	tys Tie Pi	
ttt gga gtg gg		1 2	-		
Phe Gly Val Gl 115	у пуз пец	120	ш мта сув	125	. Tyr var

		_		_		_	_				ccg Pro						432
]			_		_		_	_		_	ccg Pro 155	_	_				480
											gaa Glu						528
											aac Asn						576
		_	_							_	gtc Val						624
	_	~					_	_			ttt Phe	_	_				672
(	999 31y 225	acg Thr	agc Ser	cac His	tac Tyr	gtg Val 230	gtg Val	ggt Gly	gtg Val	gtg Val	gca Ala 235	gcg Ala	gcc Ala	gcc Ala	gtt Val	tta Leu 240	720
-					_		-				agg Arg			tga			762
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		)> 16 Ser		Gly	Ile 5	Thr	Leu	Thr	Leu	Leu 10	Leu	Ala	Thr	Leu	Ala 15	Thr	
1		Arg	Cys	Ala 20		Gln	Thr	His	Tyr 25		Ala	Val	Pro	Val 30	His	Ser	
7	Гhr	Ala	Ser 35		Gly	Cys	Val	Leu 40		Thr	Pro	His	Asp		Leu	Ile	
1	Jal	Thr 50		Gln	Lys	Gln	Glu 55		Pro	Ser	Pro	Val 60		Val	Ala	Thr	
7	Fyr 65		Ser	Glu	Ala	Gly 70		Val	Val	Gln	Pro 75		Phe	Ala	Gly	Arg 80	
7		Asp	Ile	Pro	Glu 85	His	Lys	Leu	Thr	Arg 90	Thr	Thr	Leu	Lys	Phe 95	Phe	
1	Asn	Ala	Thr	Leu 100	Glu	Asp	Glu	Gly	Cys 105	Tyr	Leu	Cys	Ile	Phe 110	Asn	Ala	
J	?he	Gly	Val 115	Gly	Lys	Leu	Ser	Gly 120	Thr	Ala	Cys	Leu	Thr 125	Val	Tyr	Val	
I	?ro	Leu 130	Ser	Met	Ser	Val	Thr 135	Phe	Tyr	Pro	Pro	Ile 140	Asn	Pro	Thr	Gln	
	Leu L45	Val	Cys	Arg	Ala	Glu 150	Ala	Ser	Pro	Ala	Pro 155	Ser	Val	Asn	Trp	Thr 160	
		Val	Pro	Pro	Glu	Leu	Cys	Ser	Glu	Pro	Glu	Val	Phe	Pro	Arg	Pro	

165 170 175 Asn Gly Thr Thr Leu Val Val Gly Arg Cys Asn Val Thr Ser Val Asp 180 185 Pro Glu Asp Leu Glu Asn Ala Thr Cys Leu Val Thr His Ile Gly Gly 200 205 195 Leu Ala Ala Arg Pro Leu Asp Pro Val Phe Ser Asp Pro Leu Glu 220 215 Gly Thr Ser His Tyr Val Val Gly Val Val Ala Ala Ala Val Leu 230 225 Gly Ile Phe Leu Thr Gly Val Phe Leu Tyr Arg Ser Met 250 245

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Gly Asn Ala Leu Val Leu Tyr Ile Phe Phe Lys Phe Lys Ala Leu Ala
65 70 75 80

55

aac tot gtg gat gta otg atg got ggg ttg tgt tgt aac too otg ttt 288 Asn Ser Val Asp Val Leu Met Ala Gly Leu Cys Cys Asn Ser Leu Phe 85 90 95

ctg tgc gcg tcg ttt ttg ttc agc tgg ctg ctg tac gtc gcg cca cag
Leu Cys Ala Ser Phe Leu Phe Ser Trp Leu Leu Tyr Val Ala Pro Gln
100 105 110

atg ctc acg tcc gcg acg tgc aag gtg gaa atc ttt ttc ttt tac ctg 384
Met Leu Thr Ser Ala Thr Cys Lys Val Glu Ile Phe Phe Phe Tyr Leu
115 120 125

tac acg tac ttt ggc gtg tac att gtg gtg tgt atc agc ctt atc agg 432
Tyr Thr Tyr Phe Gly Val Tyr Ile Val Val Cys Ile Ser Leu Ile Arg
130 135 140

tgc ctg tta gtt gtg ttt tcc cgc cgc ccg tgg gtc aag cac ggg gcc 480

Cys 145	Leu	Leu	Val	Val	Phe 150	Ser	Arg	Arg	Pro	Trp 155	Val	Lys	His	Gly	Ala 160	
			ctc Leu	_		~							_		-	528
	_		gcg Ala 180					_		-	_					576
			ata Ile	_		_	_	_	000	_	_		_			624
_	_	_	atc Ile	_			_			-			_	_	-	672
		_	atg Met	_						_		_	_	_		720
_	_	_	ctg Leu	_	_	_		_	_			_			_	768
			ctg Leu 260					_					_	_		816
		_	acc Thr	_	_					_	_	_	_	_		864
		-	gtg Val		_		_	_			_					912
_	_		agc Ser					_								960
_			aga Arg		_		_							_		1008
			ggt Gly 340	_		tga										1029

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<211> 342

<212> PRT

<213> Macaca mulatta rhadinovirus 17577

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a J	ca Thr	cgc Arg 50	ccc Pro	tct Ser	gat Asp	aac Asn	ttt Phe 55	gac Asp	aac Asn	gac Asp	gat Asp	gac Asp 60	gac Asp	cca Pro	gcg Ala	ctg Leu	192
Ç	ggc Bly 65	gtt Val	atc Ile	tgg Trp	cat His	ctt Leu 70	ctg Leu	gcg Ala	cct Pro	ctg Leu	gtt Val 75	aat Asn	tat Tyr	gca Ala	cct Pro	ctg Leu 80	240
Ç	gaa Glu	act Thr	cgg Arg	tcg Ser	gcg Ala 85	cac His	ctc Leu	cag Gln	ggc Gly	cat His 90	cat His	act Thr	ata Ile	tcc Ser	ctg Leu 95	ccc Pro	288
ţ	tat Tyr	ggc Gly	cca Pro	gac Asp 100	ctg Leu	atg Met	cgc Arg	caa Gln	cct Pro 105	acc Thr	aca Thr	aga Arg	tct Ser	agc Ser 110	gaa Glu	ata Ile	336
Ş	gtg /al	cag Gln	tgc Cys 115	ctt Leu	aga Arg	gac Asp	agc Ser	ggc Gly 120	ctc Leu	gat Asp	aga Arg	acg Thr	ttg Leu 125	cgg Arg	tta Leu	gag Glu	384
Š	gtg /al	ggc Gly 130	aga Arg	cat His	ctg Leu	agc Ser	tgc Cys 135	cag Gln	acg Thr	aga Arg	cgg Arg	ttt Phe 140	gtc Val	gcc Ala	gat Asp	cgg Arg	432
7	gta Val 145	ccc Pro	ccg Pro	ggc Gly	acc Thr	ttg Leu 150	gcc Ala	gcc Ala	ctg Leu	aca Thr	ctt Leu 155	ggc Gly	aca Thr	cta Leu	gta Val	gaa Glu 160	480
5	tat Fyr	gat Asp	gtg Val	cgc Arg	gtg Val 165	cag Gln	cgc Arg	cag Gln	ctc Leu	ccg Pro 170	gtg Val	aca Thr	ttg Leu	çaa Gln	tcc Ser 175	acc Thr	528
į	gcc Ala	tgg Trp	aga Arg	ccg Pro 180	ttg Leu	ccc Pro	gag Glu	aga Arg	gac Asp 185	cca Pro	ata Ile	tgc Cys	gcc Ala	gcg Ala 190	gtg Val	atg Met	576
]	ctc Leu	ccg Pro	tta Leu 195	caa Gln	cgg Arg	aac Asn	ata Ile	tta Leu 200	ccg Pro	ctg Leu	gcc Ala	gtg Val	cag Gln 205	gcc Ala	tcc Ser	aac Asn	624
(	ggc Gly	aac Asn 210	agc Ser	tat Tyr	acg Thr	gtg Val	tcc Ser 215	aga Arg	tac Tyr	gcc Ala	gtc Val	atg Met 220	gcc Ala	cgc Arg	agg Arg	agc Ser	672
,	tac Tyr 225	agc Ser	tgc Cys	gtt Val	ttc Phe	cag Gln 230	cgc Arg	ctc Leu	ccg Pro	tgc Cys	gaa Glu 235	aac Asn	gta Val	acc Thr	cac His	ata Ile 240	720
	gct Ala	gac Asp	tca Ser	ttt Phe	aca Thr 245	cac His	ctg Leu	cac His	agc Ser	gcc Ala 250	att Ile	cag Gln	aca Thr	ggt Gly	gca Ala 255	ggt Gly	768
į	gcg Ala	ctg Leu	caa Gln	aac Asn 260	att Ile	ctg Leu	ttc Phe	cat His	gcc Ala 265	acg Thr	ctg Leu	ctg Leu	ccc Pro	999 Gly 270	Gly	gaa Glu	816

ato Ile	aga Arg	tcg Ser 275	gcc Ala	ctg Leu	tgt Cys	gga Gly	ttt Phe 280	tac Tyr	gcc Ala	act Thr	acg Thr	ccg Pro 285	tca Ser	gtg Val	ggc Gly	864
gca Ala	ttt Phe 290	tct Ser	cgc Arg	gca Ala	cgc Arg	cac His 295	aga Arg	gct Ala	att Ile	aac Asn	aca Thr 300	aca Thr	gcg Ala	aca Thr	ctc Leu	912
cac His	tgc Cys	cag Gln	cag Gln	ctg Leu	gcg Ala 310	cgc Arg	acc Thr	ggc Gly	acg Thr	cct Pro 315	gtc Val	ctc Leu	ggt Gly	ggc Gly	ttt Phe 320	960
ct! Le:	aaa Lys	acc Thr	gtc Val	cac His 325	agc Ser	gcc Ala	acc Thr	acc Thr	agc Ser 330	gag Glu	gcg Ala	aac Asn	gtt Val	att Ile 335	acc Thr	1008
acc Th:	aca Thr	tcg Ser	ctg Leu 340	tta Leu	tcg Ser	tgc Cys	gtg Val	cct Pro 345	caa Gln	gca Ala	tac Tyr	aca Thr	ttc Phe 350	ctc Leu	agg Arg	1056
ag Ar	g tct g Ser	tta Leu 355	ttc Phe	agt Ser	cag Gln	cct Pro	atc Ile 360	atc Ile	tgt Cys	ctt Leu	gly	tct Ser 365	ttt Phe	gaa Glu	ccc Pro	1104
gt Va	gac l Asp 370	ggc Gly	gat Asp	ggc Gly	aac Asn	cag Gln 375	cgc Arg	tcg Ser	ctt Leu	tac Tyr	ctg Leu 380	ggg Gly	agc Ser	gcc Ala	gca Ala	1152
99 G1 38	t att y Ile	acc Thr	cgc Arg	atc Ile	acc Thr 390	caa Gln	acg Thr	ttg Leu	tcg Ser	ctg Leu 395	gct Ala	tac Tyr	gag Glu	att Ile	ttg Leu 400	1200
ga Gl	a ggg u Gly	ccc Pro	cta Leu	ttt Phe 405	acc Thr	agc Ser	att Ile	aat Asn	cgc Arg 410	gcc Ala	cat His	gaa Glu	ccc Pro	gcc Ala 415	tct Ser	1248
gt Va	c atc l Ile	ggc Gly	cac His 420	ctg Leu	gga Gly	gcc Ala	ctg Leu	gtc Val 425	tcg Ser	cgg Arg	ggc Gly	ggc Gly	ctg Leu 430	cgc Arg	ctc Leu	1296
tt Ph	t gtc e Val	tct Ser 435	cag Gln	ctt Leu	cca Pro	cca Pro	acc Thr 440	att Ile	ctg Leu	agc Ser	caa Gln	ctg Leu 445	acc Thr	gcc Ala	acg Thr	1344
cc Pr	a gac o Asp 450	Ile	tca Ser	cgg Arg	gaa Glu	acc Thr 455	gtg Val	aac Asn	gac Asp	atc Ile	cta Leu 460	Val	aac Asn	aag Lys	ttt Phe	1392
ct Le 46	c aac u Asn 5	gtg Val	tct Ser	gcc Ala	tgc Cys 470	gtg Val	gtc Val	ttt Phe	gcc Ala	gtc Val 475	Leu	ccg Pro	cgc Arg	gac Asp	acg Thr 480	1440
ga Gl	g ccg u Pro	gaa Glu	ccg Pro	ggc Gly 485	ccg Pro	ttg Leu	gat Asp	gcc Ala	atc Ile 490	Arg	agg Arg	gcc Ala	gca Ala	cgc Arg 495	lle	1488
t g Cy	c gga s Gly	tgc Cys	cct Pro 500	ttc Phe	gcc Ala	gtc Val	gtt Val	ggg Gly 505	gaa Glu	acc Thr	tgc Cys	gaa Glu	gag Glu 510	Leu	gga Gly	1536
at	t cag	ttc	gtg	aac	gac	ctg	gag	ctg	tgg 283	aac	ccc	gga	gcg	tgg	g ccg	1584

IJ	Le	Gln	Phe 515	Val	Asn	Asp	Leu	Glu 520	Leu	Trp	Asn	Pro	Gly 525	Ala	Trp	Pro	
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G.	ag lu 45	cag Gln	ccc Pro	gtt Val	tcc Ser	tcc Ser 550	aac Asn	tgg Trp	ctg Leu	gtg Val	cgc Arg 555	cca Pro	gaa Glu	gaa Glu	cct Pro	gag Glu 560	1680
ga As	at sp	ggt Gly	ggc Gly	gaa Glu	cag Gln 565	gca Ala	ccc Pro	tcg Ser	ccg Pro	acc Thr 570	gac Asp	tgg Trp	ggc Gly	cta Leu	ttc Phe 575	cgc Arg	1728
Ci Le	tg eu	gcc Ala	tcc Ser	gtg Val 580	gtc Val	gat Asp	cag Gln	ctt Leu	ctg Leu 585	cga Arg	tgt Cys	ccg Pro	acg Thr	gtt Val 590	ggc Gly	agc Ser	1776
aa Ly	aa ys	gag Glu	ttt Phe 595	gtc Val	acg Thr	cga Arg	cac His	gtg Val 600	gac Asp	aga Arg	tgc Cys	tcc Ser	aac Asn 605	gga Gly	ctc Leu	gtg Val	1824
g A	ct la	cag Gln 610	cag Gln	tgc Cys	gaa Glu	gtg Val	gga Gly 615	ccc Pro	ctg Leu	ggc Gly	cgg Arg	ccg Pro 620	ctg Leu	tca Ser	gat Asp	tac Tyr	1872
H	ac is 25	att Ile	gtc Val	aac Asn	cac His	acg Thr 630	tcg Ser	gtg Val	ttt Phe	acg Thr	gac Asp 635	aga Arg	atg Met	gcg Ala	cgg Arg	gtg Val 640	1920
C:	cc ro	ata Ile	tat Tyr	cgc Arg	ccc Pro 645	cag Gln	ccg Pro	atc Ile	acc Thr	agg Arg 650	cag Gln	gac Asp	gcg Ala	acg Thr	gaa Glu 655	cgc Arg	1968
L C	tg eu	gtt Val	agc Ser	cca Pro 660	gaa Glu	acc Thr	tgg Trp	gtc Val	acc Thr 665	cag Gln	ggc Gly	agg Arg	ggc Gly	agg Arg 670	aac Asn	cgg Arg	2016
t T	gg rp	gtc Val	gga Gly 675	cag Gln	tgc Cys	gtg Val	gct Ala	tat Tyr 680	gga Gly	gaa Glu	cag Gln	gca Ala	tac Tyr 685	aag Lys	atg Met	ggc Gly	2064
a I	tc le	aac Asn 690	gcg Ala	gca Ala	Val	Gly	Ala	Arg	tac Tyr	Ala	atc Ile	tgc Cys 700	gag Glu	gcg Ala	gtc Val	acc Thr	2112
A	ac sn 05	atc Ile	atg Met	cta Leu	gcg Ala	cac His 710	gtg Val	cgg Arg	cgt Arg	cta Leu	agc Ser 715	gac Asp	atc Ile	acg Thr	ctg Leu	acg Thr 720	2160
9 A	cg la	tcg Ser	gtc Val	ggt Gly	tgg Trp 725	aac Asn	ccg Pro	gag Glu	gac Asp	gac Asp 730	cag Gln	gcc Ala	tgg Trp	ctc Leu	ctg Leu 735	cag Gln	2208
C H	ac	aca Thr	ctg Leu	ttt Phe 740	gcc Ala	tgc Cys	aag Lys	gaa Glu	cta Leu 745	tgc Cys	agg Arg	gac Asp	ctg Leu	agc Ser 750	atc Ile	aac Asn	2256
t P	tc he	gcc Ala	atc Ile	acg Thr	tcg Ser	gcc Ala	ggc Gly	agc Ser	acc Thr	ccg Pro	tgc Cys	ctg Leu	tcg Ser	gaa Glu	gaa Glu	ctg Leu	2304

	755					760					765				
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